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FILE 'MEDLINE, EMBASE, BIOSIS, BIOTECHDS, SCISEARCH, HCAPLUS, NTIS, LIFESCI' ENTERED AT 14:11:05 ON 30 JAN 2004 10 S "C2GNT3" L10 S ACETYLKGLYCOSAMINE L2L3 36991 S ACETYLGLUCOSAMINE 0 S "N-ACETYLGLUCVOSAMINETRANSFERASE?" L44 DUP REM L1 (6 DUPLICATES REMOVED) L5257 S "C2GNT" L65011 S "N-ACETYLGLUCOSAMINYLTRANSFERASE?" L7 L8 5065 S L6 OR L7 6346882 S CLON? OR EXPRESS? OR RECOMBINANT L9 L10 233 S L6 AND L9 L11 146 S HUMAN AND L10 L1243 DUP REM L11 (103 DUPLICATES REMOVED) E SCHWIENTEK T/AU L13 116 S E3-E4 E CLAUSEN H/AU L14 · 804 S E3 L15 868 S L13 OR L14 L16 6 S L10 AND L15 1 DUP REM L16 (5 DUPLICATES REMOVED) L17

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FILE 'LIFESCI' ENTERED AT 14:11:05 ON 30 JAN 2004 COPYRIGHT (C) 2004 Cambridge Scientific Abstracts (CSA)

=> s "c2GnT3"

L1 10 "C2GNT3"

=> s acetylkglycosamine

L2 0 ACETYLKGLYCOSAMINE

=> s acetylglucosamine

L3 36991 ACETYLGLUCOSAMINE

=> s "N-acetylglucvosaminetransferase?"

L4 0 "N-ACETYLGLUCVOSAMINETRANSFERASE?"

=> dup rem 11

PROCESSING COMPLETED FOR L1

L5 4 DUP REM L1 (6 DUPLICATES REMOVED)

=> d 1-4 ibib ab

L5 ANSWER 1 OF 4 BIOSIS COPYRIGHT 2004 BIOLOGICAL ABSTRACTS INC. on STN

ACCESSION NUMBER: 2003:537205 BIOSIS DOCUMENT NUMBER: PREV200300537549

TITLE: UDP-N-acetylglucosamine: galactose-beta1,

3-N-acetylgalactosamine-alpha-R/(GlcNAc to GalNAc) beta1,6-N-acetylglucosaminyltransferase, C2GnT3.

AUTHOR(S): Schwientek, Tilo [Inventor, Reprint Author]; Clausen,

Henrik [Inventor]

CORPORATE SOURCE: Bronshoj, Denmark

ASSIGNEE: Glycozym Aps, Holte, Denmark

PATENT INFORMATION: US 6635461 October 21, 2003

SOURCE:

Official Gazette of the United States Patent and Trademark

Office Patents, (Oct 21 2003) Vol. 1275, No. 3. http://www.uspto.gov/web/menu/patdata.html. e-file.

ISSN: 0098-1133 (ISSN print).

DOCUMENT TYPE: LANGUAGE:

Patent English

ENTRY DATE:

Entered STN: 12 Nov 2003

Last Updated on STN: 12 Nov 2003

À novel gene defining a novel human UDP-GlcNAc: Galbeta1-3 GalNAcalpha beta1,6GlcNAc-transferase, termed C2GnT3, with unique enzymatic properties is disclosed. The enzymatic activity of C2GnT3 is shown to be distinct from that of previously identified enzymes of this gene family. The invention discloses isolated DNA molecules and DNA constructs encoding C2GnT3 and derivatives thereof by way of amino acid deletion, substitution or insertion exhibiting C2GnT3 activity, as well as cloning and expression vectors including such DNA, cells transfected with the vectors, and recombinant methods for providing C2GnT3. The enzyme C2GnT3 and C2GnT3-active derivatives thereof are disclosed, in particular soluble derivatives comprising the catalytically active domain of C2GnT3. Further, the invention discloses methods of obtaining 1,6-N-acetylglucosaminyl glycosylated saccharides, glycopeptides or glycoproteins by use of an enzymically active C2GnT3 protein or fusion protein thereof or by using cells stably transfected with a vector including DNA encoding an enzymatically active C2GnT3 protein as an expression system for recombinant production of such glycopeptides or glycoproteins. Methods are disclosed for the identification of agents with the ability to inhibit or stimulate the biological activity of C2GnT3. Furthermore, methods of using C2GnT3 in the structure-based design of inhibitors or stimulators thereof are also disclosed in the invention. Also a method for the identification of DNA sequence variations in the C2GnT3 gene by isolating DNA from a patient, amplifying C2GnT3-coding exons by PCR, and detecting the presence of DNA sequence variation, are disclosed.

ANSWER 2 OF 4 BIOTECHDS COPYRIGHT 2004 THOMSON DERWENT/ISI on STN ACCESSION NUMBER: 2001-08448 BIOTECHDS

TITLE:

New C2GnT3 polypeptides and nucleic acids encoding the polypeptides useful for treating conditions mediated by a

C2GnT3 polypeptide, e.g. thymus-related disorders,

cancers, tumors, immunosuppression;

vector-mediated gene transfer, expression in bacterium, yeast, avian, mammal, CHO or Sf9 cell, DNA probe and antibody for recombinant protein production, drug

screening and gene therapy

AUTHOR:

PATENT INFO:

Schwientek T; Clausen H PATENT ASSIGNEE: Schwientek T; Clausen H

LOCATION:

Bronshoj, Denmark; Holte, Denmark. WO 2001014535 1 Mar 2001

PRIORITY INFO:

APPLICATION INFO: WO 2000-DK469 24 Aug 2000 US 1999-150488 24 Aug 1999

DOCUMENT TYPE: LANGUAGE:

Patent English

OTHER SOURCE:

WPI: 2001-226615 [23]

An isolated DNA (I, having specified 1,362 bp sequence) encoding a UDP-N-acetyl-glucosamine:galactose-beta-1,3-N-acetylgalactosamine-alpha-Rbeta-1-6-N-acetylglucosaminyltransferase (C2GnT3), or its fragment, is claimed. Also claimed are: a DNA vector (II) containing a DNA sequence encoding C2GnT3 or its fragments; a cell (III, e.g. bacterium, yeast, avian, mammal, CHO or Spodoptera frugiperda Sf9 cell) containing (II); producing C2GnT3 proteins; screening one or more agents for the ability to inhibit or stimulate C2GnT3 enzymatic activity in a cell-free or cell-based assay; identifying DNA sequence variations in the C2GnT3 gene; a C2GnT3

protein (IV) having a specified 453 amino acid protein sequence; an antibody specific against an epitope; a DNA probe containing a sequence encoding (IV); diagnosing and monitoring conditions mediated by (IV); identifying a substance which associates with (IV); evaluating a compound which modulates (IV) activity; gene therapy directed at the thymus; and preparing an oligosaccharide. (I) or (IV) is useful in the preparation of compositions for treating conditions mediated by (IV), particularly a thymus-related disorder. (96pp)

L5 ANSWER 3 OF 4 MEDLINE on STN DUPLICATE 2

ACCESSION NUMBER: 2000219156 MEDLINE

DOCUMENT NUMBER: 20219156 PubMed ID: 10753916

TITLE: Control of O-glycan branch formation. Molecular cloning and

characterization of a novel thymus-associated core 2 beta1,

6-n-acetylglucosaminyltransferase.

AUTHOR: Schwientek T; Yeh J C; Levery S B; Keck B; Merkx G; van

Kessel A G; Fukuda M; Clausen H

CORPORATE SOURCE: School of Dentistry, University of Copenhagen, Norre Alle

20, 2200 Copenhagen N, Denmark.. tsc@odont.ku.dk

CONTRACT NUMBER: 5 P41 RR05351 (NCRR)

SOURCE: JOURNAL OF BIOLOGICAL CHEMISTRY, (2000 Apr 14) 275 (15)

11106-13.

Journal code: 2985121R. ISSN: 0021-9258.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals OTHER SOURCE: GENBANK-AF132035

ENTRY MONTH: 200005

ENTRY DATE: Entered STN: 20000518

Last Updated on STN: 20000518 Entered Medline: 20000505

AB Core 2 O-glycan branching catalyzed by UDP-N-acetyl-alpha-D-glucosamine: acceptor beta1, 6-N-acetylglucosaminyltransferases (beta6GlcNAc-Ts) is an important step in mucin-type biosynthesis. Core 2 complex-type O-glycans are involved in selectin-mediated adhesion events, and O-glycan branching appears to be highly regulated. Two homologous beta6GlcNAc-Ts functioning in O-glycan branching have previously been characterized, and here we report a third homologous beta6GlcNAc-T designated C2GnT3. C2GnT3 was identified by BLAST analysis of human genome survey sequences. The catalytic activity of C2GnT3 was evaluated by in vitro analysis of a secreted form of the protein expressed in insect cells. The results revealed exclusive core 2 beta6GlcNAc-T activity. The product formed with core 1-para-nitrophenyl was confirmed by (1)H NMR to be core 2-para-nitrophenyl. In vivo analysis of the function of C2GnT3 by coexpression of leukosialin (CD43) and a full coding construct of C2GnT3 in Chinese hamster ovary cells confirmed the core 2 activity and failed to reveal I activity. The C2GnT3 gene was located to 5q12, and the coding region was contained in a single exon. Northern analysis revealed selectively high levels of a 5.5-kilobase C2GnT3 transcript in thymus with only low levels in other organs. The unique expression pattern of C2GnT3 suggests that this enzyme serves a specific function different from other members of the beta6GlcNAc-T gene family.

L5 ANSWER 4 OF 4 BIOSIS COPYRIGHT 2004 BIOLOGICAL ABSTRACTS INC. on STN

ACCESSION NUMBER: 2001:93139 BIOSIS DOCUMENT NUMBER: PREV200100093139

AUTHOR (S):

TITLE: Molecular cloning and characterization of a novel thymus

associated core 2 beta1,6-N-acetylglucosaminyltransferase. Schwientek, Tilo [Reprint author]; Levery, Steven B.; Yeh, Jiunn-Chern; Keck, Birgit [Reprint author]; Merkx, Gerard;

van Kessel, Ad Geurts; Fukuda, Minoru; Clausen, Henrik

CORPORATE SOURCE: School of Dentistry, University of Copenhagen, Norre Alle

20, 2200, Copenhagen N, Denmark SOURCE: Glycoconjugate Journal, (January-February, 2000) Vol. 17, No. 1-2, pp. 49. print. Meeting Info.: Second International Glycosyltransferase Symposium. Toronto, Ontario, Canada. May 12-14, 2000. ISSN: 0282-0080. DOCUMENT TYPE: Conference; (Meeting) Conference; Abstract; (Meeting Abstract) LANGUAGE: English ENTRY DATE: Entered STN: 21 Feb 2001 Last Updated on STN: 12 Feb 2002 => s "c2GnT" L6 257 "C2GNT" => d his (FILE 'HOME' ENTERED AT 14:09:33 ON 30 JAN 2004) FILE 'MEDLINE, EMBASE, BIOSIS, BIOTECHDS, SCISEARCH, HCAPLUS, NTIS, LIFESCI' ENTERED AT 14:11:05 ON 30 JAN 2004 10 S "C2GNT3" L10 S ACETYLKGLYCOSAMINE L2L336991 S ACETYLGLUCOSAMINE 0 S "N-ACETYLGLUCVOSAMINETRANSFERASE?" L4L54 DUP REM L1 (6 DUPLICATES REMOVED) 257 S "C2GNT" L6=> s "N-acetylglucosaminyltransferase?" 5011 "N-ACETYLGLUCOSAMINYLTRANSFERASE?" 1.7 => s 16 or 17 5065 L6 OR L7 1.8 => s clon? or express? or recombinant · 5 FILES SEARCHED... 6346882 CLON? OR EXPRESS? OR RECOMBINANT => s 16 and 19 233 L6 AND L9 L10 => s human and l10 T.11 146 HUMAN AND L10 => dup rem 111 PROCESSING COMPLETED FOR L11 43 DUP REM L11 (103 DUPLICATES REMOVED) => d 1-43 ibib ab L12 ANSWER 1 OF 43 SCISEARCH COPYRIGHT 2004 THOMSON ISI on STN ACCESSION NUMBER: 2003:1003351 SCISEARCH THE GENUINE ARTICLE: 741HH Highly conserved cysteines of mouse core 2 beta TITLE:

1,6-N-acetylglucosaminyltransferase I form a network of disulfide bonds and include a thiol that affects enzyme

activity

AUTHOR: Yen T Y; Macher B A; Bryson S; Chang X Q; Tvaroska I; Tse

R; Takeshita S; Lew A M; Datti A (Reprint)

CORPORATE SOURCE: GlycoDesign Inc, 480 Univ Ave, Ste 400, Toronto, ON M5G

1V2, Canada (Reprint); GlycoDesign Inc, Toronto, ON M5G 1V2, Canada; San Francisco State Univ, Dept Chem &

Biochem, San Francisco, CA 94132 USA; Seikagaku Corp, Cent

Res Labs, Tokyo 2070021, Japan

COUNTRY OF AUTHOR:

Canada; USA; Japan

SOURCE:

JOURNAL OF BIOLOGICAL CHEMISTRY, (14 NOV 2003) Vol. 278,

No. 46, pp. 45864-45881.

Publisher: AMER SOC BIOCHEMISTRY MOLECULAR BIOLOGY INC,

9650 ROCKVILLE PIKE, BETHESDA, MD 20814-3996 USA.

ISSN: 0021-9258. Article; Journal

DOCUMENT TYPE:

English

LANGUAGE:

Engli

REFERENCE COUNT:

ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS

Core 2 beta1,6-N-acetylglucosaminyltransferase I (C2GnT-I) plays a pivotal role in the biosynthesis of mucin-type O-glycans that serve as ligands in cell adhesion. To elucidate the three-dimensional structure of the enzyme for use in computer-aided design of therapeutically relevant enzyme inhibitors, we investigated the participation of cysteine residues in disulfide linkages in a purified murine recombinant enzyme. The pattern of free and disulfide-bonded Cys residues was determined by liquid chromatography/electrospray ionization tandem mass spectrometry in the absence and presence of dithiothreitol. Of nine highly conserved Cys residues, under both conditions, one (Cys(217)) is a free thiol, and eight are engaged in disulfide bonds, with pairs formed between Cys(59) - Cys(413), Cys(100) - Cys(172), Cys(151) - Cys(199), and Cys(372)-Cys(381). The only non-conserved residue within the beta1,6-N-acetylglucosaminyltransferase family, Cys(235), is also a free thiol in the presence of dithiothreitol; however, in the absence of reductant, Cys(235) forms an intermolecular disulfide linkage. Biochemical studies performed with thiol-reactive agents demonstrated that at least one free cysteine affects enzyme activity and is proximal to the UDP-GlcNAc binding site. A Cys(217)-->Ser mutant enzyme was insensitive to thiol reactants and displayed kinetic properties virtually identical to those of the wild-type enzyme, thereby showing that Cys(217), although not required for activity per se, represents the only thiol that causes enzyme inactivation when modified. Based on the pattern of free and disulfide-linked Cys residues, and a method of fold recognition/threading and homology modeling, we have computed a three-dimensional model for this enzyme that was refined using the T4 bacteriophage beta-

L12 ANSWER 2 OF 43 BIOSIS COPYRIGHT 2004 BIOLOGICAL ABSTRACTS INC. on STN ACCESSION NUMBER: 2004:5496 BIOSIS

ACCESSION NUMBER: DOCUMENT NUMBER:

PREV200400006400

TITLE:

Core 2 fA-1,6-N-acetylglucosaminyltransferase (

C2GnT) expression in human

prostate cancer: A predictor for non-organ confined disease

and biochemical relapse after radical prostatectomy. Ohyama, Chikara [Reprint Author]; Hagisawa, Shigeru;

Hatakeyama, Shingo [Reprint Author]; Arai, Yoichi; Fukuda,

Minoru

glucosyltransferase fold.

CORPORATE SOURCE:

Department of Urology, Akita University School of Medicine,

Akita, Japan

SOURCE:

AUTHOR (S):

Glycobiology, (November 2003) Vol. 13, No. 11, pp. 864.

print.

Meeting Info.: 8th Annual Conference of the Society for Glycobiology. San Diego, California, USA. December 03-06,

2003. Society for Glycobiology.

ISSN: 0959-6658.

DOCUMENT TYPE:

Conference; (Meeting)

Conference; Abstract; (Meeting Abstract)

LANGUAGE:

English

ENTRY DATE:

Entered STN: 17 Dec 2003

Last Updated on STN: 17 Dec 2003

L12 ANSWER 3 OF 43 SCISEARCH COPYRIGHT 2004 THOMSON ISI on STN

2003:1076263 SCISEARCH ACCESSION NUMBER:

THE GENUINE ARTICLE: 735WU

TITLE: Core 2 fA-1,6-N-acetylglucosaminyltransferase (

C2GnT) expression in human

prostate cancer: A predictor for non-organ confined

disease and biochemical relapse after radical

prostatectomy

AUTHOR: Ohyama C (Reprint); Hagisawa S; Hatakeyama S; Arai Y I;

Fukuda M

CORPORATE SOURCE: Akita Univ, Sch Med, Dept Urol, Akita 010, Japan; Tohoku

Univ, Sch Med, Dept Urol, Sendai, Miyagi 980, Japan;

Burnham Inst, Glycobiol program, La Jolla, CA 92037 USA

COUNTRY OF AUTHOR: Japan; USA

SOURCE:

GLYCOBIOLOGY, (NOV 2003) Vol. 13, No. 11, pp. 864-864. MA

143.

Publisher: OXFORD UNIV PRESS INC, JOURNALS DEPT, 2001

EVANS RD, CARY, NC 27513 USA.

ISSN: 0959-6658.

DOCUMENT TYPE:

Conference; Journal

LANGUAGE:

English

REFERENCE COUNT:

L12 ANSWER 4 OF 43 MEDLINE on STN DUPLICATE 1

ACCESSION NUMBER: DOCUMENT NUMBER:

2003248675 MEDLINE PubMed ID: 12626388

TITLE:

Multiple transcription initiation and alternative splicing

in the 5' untranslated region of the core 2 beta1-6

N-acetylglucosaminyltransferase I gene.

AUTHOR:

Falkenberg V Rebecca; Alvarez Karen; Roman Clara; Fregien

Nevis

CORPORATE SOURCE:

Department of Cell Biology and Anatomy, University of Miami

School of Medicine, Miami, FL 33176, USA.

SOURCE:

Glycobiology, (2003 Jun) 13 (6) 411-8. Journal code: 9104124. ISSN: 0959-6658.

PUB. COUNTRY:

England: United Kingdom

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

200401

ENTRY DATE:

Entered STN: 20030530

Last Updated on STN: 20040106

Entered Medline: 20040105

AB The glycosyltransferase core 2 beta1,6 N-acetylglucosaminyltransferase I (C2GnT I) plays an important regulatory role in the synthesis of biologically significant oligosaccharide structures. This gene is expressed in a variety of cell types, including lymphocytes and mucin-producing cells. The expression pattern of this gene suggests a complex system of regulation. To investigate the molecular regulation of this gene and locate potential promoter elements, rapid amplification of cDNA ends (RACE) analysis was used to determine the 5' ends of the C2GnT I mRNAs from a number of tissues. These experiments identified five C2GnT I mRNAs that are different in their 5' untranslated regions. The RACE cDNAs had four different 5' terminal sequences (exons A, B, D, and E'), suggesting four transcription initiation sites. One mRNA form was the result of alternative exon (exon C) utilization. These exons are spread across 60 kb of DNA on human chromosome 9, and all splice to the exon (exon F) that contains the C2GnT I coding region. Reverse transcription polymerase chain reaction experiments using primers specific for each of the four 5' end exon sequences revealed that the 5' terminal exons are differentially expressed, suggesting tissue specificity for the different 5' untranslated regions. These findings are consistent with the presence of multiple tissue-specific promoters for the C2GnT I

L12 ANSWER 5 OF 43

MEDLINE on STN

DUPLICATE 2

ACCESSION NUMBER: DOCUMENT NUMBER:

2003227343

IN-PROCESS

PubMed ID: 12626393

TITLE:

Purification and cDNA cloning of

UDP-GlcNAc:GlcNAcbeta1-3Galbeta1-4Glc(NAc)-R [GlcNAc to Gal]beta1,6N-acetylglucosaminyltransferase from rat small intestine: a major carrier of dIGnT activity in rat small

intestine.

AUTHOR:

Korekane Hiroaki; Taguchi Tomohiko; Sakamoto Yoshihiro; Honke Koichi; Dohmae Naoshi; Salminen Heidi; Toivonen Suvi; Helin Jari; Takio Koji; Renkonen Ossi; Taniguchi Naoyuki

CORPORATE SOURCE:

Department of Biochemistry, Osaka University Medical School/graduate School of Medicine, 2-2 Yamadaoka, Suita,

Osaka 565-0871, Japan.

SOURCE:

Glycobiology, (2003 May) 13 (5) 387-400. Journal code: 9104124. ISSN: 0959-6658.

PUB. COUNTRY:

England: United Kingdom

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

IN-PROCESS; NONINDEXED; Priority Journals

OTHER SOURCE:

GENBANK-AB098520

ENTRY DATE:

Entered STN: 20030517

Last Updated on STN: 20031217

A rat intestinal beta1,6N-acetylglucosaminyltransferase (beta1-6GnT) AB responsible for the formation of the beta1,6-branched poly-Nacetyllactosamine structure has been purified to apparent homogeneity by successive column chromatographic procedures using an assay wherein pyridylaminated lacto- N-triose II (GlcNAcbeta1-3Galbeta1-4Glc-PA) was used as an acceptor substrate and the reaction product was GlcNAcbeta1-3(GlcNAcbeta1-6)Galbeta1-4Glc-PA. The purified enzyme catalyzed the conversion of the polylactosamine acceptor GlcNAcbeta1-3'LacNAc into GlcNAcbeta1-3'(GlcNAcbeta1-6') LacNAc (dIGnT activity), but it could not transfer GlcNAc to LacNAcbeta1-3'LacNAc (cIGnT activity). This enzyme could also convert mucin core 1 and core 3 analogs, Galbeta1-3GalNAcalpha1-O-paranitrophenyl (pNP) and GlcNAcbeta1-3GalNAcalpha1-O-pNP, into Galbeta1-3(GlcNAcbeta1-6) GalNAcalpha1-O-pNP (C2GnT activity) and GlcNAcbeta1-3(GlcNAcbetal-6)GalNAcalphal-O-pNP (C4GnT activity), respectively. on the partial amino acid sequences of the purified protein, the cDNA encoding this enzyme was cloned. The COS-1 cells transiently transfected with this cDNA had high dI/C2/C4GnT activities in a ratio of 0.34:1.00:0.90, compared with non- or mock-transfected cells. The primary structure shows a significant homology with human and viral mucin-type core 2 betal-6GnTs (${\tt C2GnT-Ms}$), indicating that this enzyme is the rat ortholog of **human** and viral ${\tt C2GnT}$ This is the first identification and purification of this enzyme as a major carrier of dIGnT activity in the small intestine. This rat ortholog should mostly be responsible for making distal I-branch structures on poly-N-acetyllactosamine sequences in this tissue, as well as making mucin core 2 and core 4 structures, given that it also has high C2/C4GnT activities.

L12 ANSWER 6 OF 43

MEDLINE on STN

DUPLICATE 3

ACCESSION NUMBER:

2003293624 MEDLINE

DOCUMENT NUMBER:

22705153 PubMed ID: 12600830

Mucin biosynthesis: epidermal growth factor downregulates

core 2 enzymes in a human airway adenocarcinoma

cell line.

AUTHOR:

Beum Paul V; Bastola Dhundy R; Cheng Pi-Wan

CORPORATE SOURCE:

Department of Biochemistry and Molecular Biology, University of Nebraska Medical Center, Omaha, Nebraska

68198-4525, USA.

CONTRACT NUMBER:

R01 HL48282 (NHLBI)

SOURCE:

AMERICAN JOURNAL OF RESPIRATORY CELL AND MOLECULAR BIOLOGY,

(2003 Jul) 29 (1) 48-56.

Journal code: 8917225. ISSN: 1044-1549.

PUB. COUNTRY:

United States

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

200308

ENTRY DATE:

Entered STN: 20030625

Last Updated on STN: 20030806 Entered Medline: 20030805

Enzymes which exhibit core 2 beta1,6 N-acetylglucosaminyltransferase (AB C2GnT) activity play important roles in physiologic processes including the inflammatory response and immune system function, and C2GnT activity is regulated during processes, such as T cell activation and cellular differentiation. In this study, we have examined the regulation of C2GnT activity in the H292 airway epithelial cell line by epidermal growth factor (EGF), which has been previously shown to upregulate expression of the airway mucin MUC5AC in this cell line. We found that EGF suppressed C2GnT activity in a time- and dose-dependent fashion, and also suppressed core 4 beta1,6 N-acetylglucosaminyltransferase (C4GnT) activity. Consistent with the suppression of C4GnT activity, Northern blotting results showed that EGF preferentially inhibited the M isoform of C2GnT, which forms core 2, core 4, and blood group I betal, 6 branched carbohydrate structures, while the L isoform, which forms only the core 2 structure, was only modestly affected. Furthermore, EGF treatment resulted in a shift in the carbohydrate structure of FLAG-tagged MUC1 expressed in the cells from core 2-based toward core 1-based structures, consistent with the inhibitory effects of EGF on C2GnT. Transforming growth factor alpha mimicked the effect of EGF on C2GnT, implicating the EGF receptor (EGF-R) in C2GnT suppression, and the EGF-R tyrosine kinase inhibitor AG1478 blocked C2GnT suppression, confirming the role of EGF-R in the inhibition of C2GnT expression. Also, PD98059, a specific inhibitor of mitogen-activated protein kinase/extracellular signal-regulated kinase kinase (MEK)1/2 in the Ras-mitogen-activated protein kinase pathway, completely blocked the EGF suppressive effect, suggesting possible involvement of the Ras-mitogen-activated protein kinase pathway in EGF-mediated downregulation of C2GnT. The results of this study suggest that exposure of airway cells to EGF may result in remodeling of mucin carbohydrate structure, potentially altering the biological properties of the cells.

L12 ANSWER 7 OF 43

NTIS COPYRIGHT 2004 NTIS on STN

ACCESSION NUMBER: NTIS ORDER NUMBER:

2003(19):00303 ADA413351/XAB

TITLE:

AUTHOR:

Elucidation of a Novel Cell Death Mechanism in Prostate Epithelial Cells. Annual rept. 19 Nov 2001-18 Nov 2002.

Baum, L. G.

CORPORATE SOURCE:

California Univ., Los Angeles. (005420000 072250)

NUMBER OF REPORT:

ADA413351/XAB

-

21p; Dec 2002 DAMD17-02-1-0022

NUMBER OF CONTRACT: CONTROLLED TERM:

Report

COUNTRY:

United States

LANGUAGE:

English

NOTES:

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22161, USA.

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GRA&I0319

Tumor cell resistance to apoptosis is a major obstacle to effective therapy of prostate cancer. We have found that the androgen dependent prostate cancer cell line LNCaP is sensitive to apoptosis induced by galectin-1, an endogenous human lectin that is abundant in prostate stroma. In contrast, androgen independent LNCaP, DU145 and PC3 cells are resistant to galectin-1 induced death and actually synthesize galectin-1 and export it to the cell surface. Galectin-1 binds to saccharide ligands on susceptible LNCaP cells to trigger cell death. Susceptibility to galectin-1 appears to depend on the presence of a specific class of cell surface glycans, the O-linked glycans on glycoproteins; in contrast, N-glycans are not required for galectin-1 induced LNCaP cell death. Resistance to galectin-1 induced death correlates with markedly decreased expression of a specific glycosyltransferase, the C2GnT, which creates saccharide ligands on O-glycans that are recognized by galectin-1. The C2GnT enzyme also regulates susceptibility of T cells to galectin-1 induced death, indicating that a common glycosylation pathway may control cell death in epithelial and lymphoid cells. Identification of a mechanism that enhances galectin-1 prostate cancer cell death may allow novel therapeutic approaches to manipulate tumor cell glycosylation to overcome tumor cell resistance to apoptosis.

L12 ANSWER 8 OF 43 MEDLINE on STN DUPLICATE 4

ACCESSION NUMBER:

2002696058

MEDLINE

DOCUMENT NUMBER:

PubMed ID: 12359718 22344688

TITLE:

The alpha (1,3)-fucosyltransferase Fuc-TIV, but not Fuc-TVII, generates sialyl Lewis X-like epitopes

preferentially on glycolipids.

AUTHOR:

Huang Min-Chuan; Laskowska Anna; Vestweber Dietmar; Wild

Martin K

CORPORATE SOURCE:

Institute of Cell Biology, Center for Molecular Biology of

Inflammation, University of Munster and the

Max-Planck-Institute of Vascular Biology, D-48149 Munster,

SOURCE:

JOURNAL OF BIOLOGICAL CHEMISTRY, (2002 Dec 6) 277 (49)

47786-95.

Journal code: 2985121R. ISSN: 0021-9258.

PUB. COUNTRY:

United States

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

200302

ENTRY DATE:

Entered STN: 20021217

Last Updated on STN: 20030205

Entered Medline: 20030204

AΒ Fuc-TIV and Fuc-TVII are the two alpha(1, 3)-fucosyltransferases in myeloid cells responsible for the biosynthesis of sialyl Lewis X (sLe(x)), the minimal ligand structure for the selectins. We have compared the ability of Fuc-TIV and Fuc-TVII to generate sLe(x)-like epitopes in transfected Chinese hamster ovary (CHO) - Pro(-) 5 cells expressing the P-selectin glycoprotein ligand-1 and the core-2 branching enzyme C2GnT. We found that mouse Fuc-TIV and Fuc-TVII can generate similar levels of cell surface sLe(x). Surprisingly however, Fuc-TIV-generated sLe(x) was resistant to proteinase K and trypsin treatment and could be removed from cells by delipidation with chloroform/methanol, whereas 80-90% of Fuc-TVII-generated sLe(x) was protease-sensitive, and most of it resistant to delipidation. Despite similar levels of sLe(x) on the cell surface, Fuc-TVII transfectants adhered to immobilized E-selectin-IgG under static and flow conditions better than Fuc-TIV transfectants. Binding was mainly protease sensitive, indicating that glycoproteins were more efficient ligands than glycolipids. In summary, we conclude that the two fucosyltransferases differ in their in vivo specificity for acceptor substrates with Fuc-TVII generating sLe(x) preferentially on glycoproteins, whereas most of the Fuc-TIV-generated sLe(x) is found on glycolipids. Interestingly, the non-catalytic portion of Fuc-TIV in a Fuc-TIV/VII chimeric enzyme mediated the specificity for glycolipid substrates.

·L12 ANSWER 9 OF 43

MEDLINE on STN

DUPLICATE 5

ACCESSION NUMBER:

2002299855

MEDLINE

DOCUMENT NUMBER:

22005981 PubMed ID: 12010808

TITLE:

The monoclonal antibody CHO-131 binds to a core 2 O-glycan

terminated with sialyl-Lewis x, which is a functional

glycan ligand for P-selectin.

AUTHOR:

Walcheck Bruce; Leppanen Anne; Cummings Richard D; Knibbs Randall N; Stoolman Lloyd M; Alexander Shelia R; Mattila

Polly E; McEver Rodger P

CORPORATE SOURCE:

Department of Veterinary PathoBiology, University Minnesota Academic Health Center, University of Minnesota, St Paul

55108, USA.. walch003@umn.edu

CONTRACT NUMBER:

AI 48075 (NIAID)

HL 65631 (NHLBI)

SOURCE:

BLOOD, (2002 Jun 1) 99 (11) 4063-9.

Journal code: 7603509. ISSN: 0006-4971.

PUB. COUNTRY: DOCUMENT TYPE: United States

Journal; Article; (JOURNAL ARTICLE) English

LANGUAGE:

FILE SEGMENT:

Abridged Index Medicus Journals; Priority Journals

ENTRY MONTH:

200207

ENTRY DATE:

Entered STN: 20020604

Last Updated on STN: 20020702

Entered Medline: 20020701 AB Core 2 O-glycans terminated with sialyl-Lewis x (sLe(X)) are functionally important oligosaccharides that endow particular macromolecules with high-affinity glycan ligands for the selectin family. To date, antibodies that recognize these structures on leukocytes have not been described. We characterize such a monoclonal antibody (mAb) here (CHO-131). The binding specificity of CHO-131 was directly examined by means of synthetic glycopeptides containing precise O-glycan structures. CHO-131 bound to sLe(X) extended from a core 2 branch (C2-O-sLe(X)), but CHO-131 demonstrated no reactivity if this oligosaccharide lacked fucose or if sLe(X) was extended from a core 1 branch. Using transfected cell lines, we found that CHO-131 binding required the functional activity of the glycosyltransferases alpha2,3-sialyltransferase, alpha1,3fucosyltransferase-VII, and core 2 beta1,6 N-acetylglucosaminyltransferase (C2GnT). The C2-O-sLe(X) motif occurs primarily on sialomucins and has been directly shown to contribute to high-affinity P-selectin glycoprotein ligand-1 binding by P-selectin. Indeed, CHO-131 staining of neutrophils was diminished following sialomucin removal by O-glycoprotease, and its reactivity with transfected hematopoietic cell lines correlated with the expression of P-selectin ligands. CHO-131 also stained a small population of lymphocytes that were primarily CD3(+), CD4(+), and CD45RO(+) and represented a subset (37.8% +/- 18.3%) of cutaneous lymphocyte-associated antigen (CLA) T cells, distinguished by the mAb HECA-452, which detects sLe(X)-related glycans. Unlike anti-sLe(X) mAbs, CHO-131 binding also indicates C2GnT activity and demonstrates that CLA T cells are heterogeneous based on the glycan structures they synthesize. These findings support evidence that differential C2GnT activity results in T-cell subsets that

L12 ANSWER 10 OF 43

MEDLINE on STN

express ligands for E-selectin, P-selectin, or both.

DUPLICATE 6

ACCESSION NUMBER:

2002450937

MEDLINE 22197242 PubMed ID: 12209829

DOCUMENT NUMBER:

TITLE: Engineering of coordinated up- and down-regulation of two

glycosyltransferases of the O-glycosylation pathway in

Chinese hamster ovary (CHO) cells.

AUTHOR: Prati Elisabetta G P; Matasci Mattia; Suter Tobias B;

Dinter Andre; Sburlati Adriana R; Bailey James E

CORPORATE SOURCE: Institute of Biotechnology, ETH Zurich, Switzerland...

prati@biotech.biol.ethz.ch

SOURCE: BIOTECHNOLOGY AND BIOENGINEERING, (2002 Sep 5) 79 (5)

580-5.

Journal code: 7502021. ISSN: 0006-3592.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200303

ENTRY DATE: Entered STN: 20020906

Last Updated on STN: 20030320 Entered Medline: 20030319

AB Production of O-linked oligosaccharides that interact with selectins to mediate cell-cell adhesion occurs in one segment of a branched glycan biosynthesis network. Prior efforts to direct the branched pathway towards selectin-binding oligosaccharides by amplifying enzymes in this branch of the network have had limited success, suggesting that metabolic engineering to simultaneously inhibit the competing pathway may also be required. We report here the partial cloning of the CMP-sialic, acid:Galbeta1,3GalNAcalpha2,3-sialyltransferase (ST3Gal I) gene from Chinese hamster ovary (CHO) cells and the simultaneous inhibition of expression of CHO cell ST3Gal I gene and overexpression of the human UDP-GlcNAc:Galbeta1,3GalNAc-R beta1,6-Nacetylglucosaminyltransferase (C2GnT) gene. A tetracycline-regulated system adjoined to tricistronic expression technology allowed "one-step" transient manipulation of multiple enzyme activities in the O-glycosylation pathway of a previously established CHO cell line already engineered to express alpha1,3fucosyltransferase VI (alpha1,3-Fuc-TVI). Tetracycline-regulated coexpression of a ST3Gal I fragment, cloned in the

antisense orientation, and of C2GnT cDNA resulted in inhibition of the ST3Gal I enzymatic activity and increase in C2GnT activity which varied depending on the extent of tetracycline reduction in the cell culture medium. This simultaneous regulated inhibition and activation of the two key enzyme activities in the O-glycosylation pathway of mammalian cells is an important addition to the metabolic engineering

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L12 ANSWER 11 OF 43 MEDLINE on STN

ACCESSION NUMBER: 2003138543 MEDLINE

DOCUMENT NUMBER: 22540114 PubMed ID: 12652802

TITLE: Glycosyltransferase genes as tumor marker.

AUTHOR: Nakayama Jun: Shimizu Fumiaki: Katsuyama Tsu

AUTHOR: Nakayama Jun; Shimizu Fumiaki; Katsuyama Tsutomu
CORPORATE SOURCE: Department of Pathology, Shinshu University School of

Medicine.

SOURCE: RINSHO BYORI. JAPANESE JOURNAL OF CLINICAL PATHOLOGY, (2002

Nov) Suppl 123 142-8. Ref: 15

Journal code: 2984781R. ISSN: 0047-1860.

PUB. COUNTRY: Japan

field.

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

General Review; (REVIEW)

(REVIEW, TUTORIAL)

LANGUAGE: Japanese

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200305

ENTRY DATE: Entered STN: 20030326

Last Updated on STN: 20030521

Entered Medline: 20030520

Core 2 beta 1,6-N-acetylglucosaminyltransferase(C2GnT) and alpha AΒ 1,4-N-acetylglucosaminyltransferase are glycosyltransferases involved in the biosynthesis of mucin type glycoprotein(O-glycan). The transcripts of C2GnT, which forms core 2-branched O-glycan(Gal beta 1-->3 (GlcNAc beta 1-->6)GalNAc alpha-->Ser/Thr), were detected approximately in 2/3 cases of patients with colorectal or lung cancers. Then, carcinoma cells expressing C2GnT mRNA were shown to significantly progress compared with those lacking the C2GnT mRNA, indicating an important role of the core 2-branched O-glycan in tumor progression. On the other hand, gland mucous cell-type mucin secreted from the normal gastric mucosa characteristically contains GlcNAc alpha 1-->4Gal beta-->R structure, and the alpha 4GnT is critical for the biosynthesis of this unique glycan. This enzyme is also detected in gastric cancer cells but not in mononuclear cell fraction of the peripheral blood. Thus, the quantitative RT-PCR method targeted to alpha 4GnT mRNA will be useful for the detection of circulating gastric cancer cells in the peripheral blood.

L12 ANSWER 12 OF 43 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER:

2001:431030 HCAPLUS

DOCUMENT NUMBER:

135:165285

TITLE:

Overexpression of sialyltransferase CMP-sialic

acid:Gal.beta.1,3GalNAc-R .alpha.6-sialyltransferase

is related to poor patient survival in human

colorectal carcinomas

AUTHOR (S):

Schneider, Frank; Kemmner, Wolfgang; Haensch,

Wolfgang; Franke, Gudrun; Gretschel, Stephan; Karsten,

Uwe; Schlag, Peter Michael

CORPORATE SOURCE:

Department for Surgery and Surgical Oncology,

Robert-Rossle-Klinik at the Max Delbruck Center for Molecular Medicine, Charite, Berlin, D-13122, Germany

SOURCE: Cancer Research (2001), 61(11), 4605-4611

CODEN: CNREA8; ISSN: 0008-5472

PUBLISHER:

American Association for Cancer Research

DOCUMENT TYPE: Journal LANGUAGE: English

Thomsen-Friedenreich (TF)-related blood group antigens, such as TF, Tn, and their sialylated variants, belong to a family of tumor-assocd. carbohydrates. The aim of the present study was to examine tumor-assocd. alterations of glycosyltransferases involved in the biosynthesis of the TF glycotope in colorectal carcinomas. To this end, glycosyltransferase expression was examd. in 40 cases of colorectal carcinoma specimens classified according to the WHO/Union International Contre Cancer guidelines and in "normal" mucosa of the same patients. Occurrence of TF glycotope was examd. by immunohistochem. with the monoclonal antibody A78-G/A7. Expression of sialyltransferases CMP-sialic acid:Gal.beta.1,3GalNAc-R .alpha.3-sialyltransferase I and II (ST3Gal-I and ST3Gal-II) and CMP-sialic acid:Gal.beta.1,3GalNAc-R .alpha.6-sialyltransferase (ST6GalNAc-II) and of core 2 .beta.1,6-N-acetylglucosaminyltransferase was detd. by reverse transcription-PCR in the same cryostat sections used for immunohistochem. Addnl., .alpha.2,3-sialyltransferase enzyme activity was studied in each of these tissues. The TF glycotope was detected in 7% of the normal mucosa, but in 57% of the carcinoma samples. Expression of .alpha.2,3-sialyltransferases ST3Gal-I, ST3Gal-II, and enzyme activity of .alpha.2,3-sialyltransferase was significantly increased (P < 0.001) in carcinoma specimens compared with normal mucosa. ST3Gal-I mRNA expression was significantly increased (P = 0.05) in cases showing invasion of lymph vessels. Expression of ST6GalNAc-II was significantly increased (P = 0.04) in cases with metastases to lymph nodes along the vascular trunk. Moreover, ST6GalNAc-II expression provides an prognostic factor for patient survival ($log\ rank$, P = 0.02). In an attempt to study the functional relevance of the glycosyltransferases for TF biosynthesis, SW480 colorectal cells were

transfected with each of the enzymes, and cell surface expression of the TF glycotope was examd. by flow cytometry. The presence of TF was not altered by transfection of the cells with either sialyltransferase ST3Gal-I or ST3Gal-II. However, successful transfection with core 2 .beta.1,6-N-acetylglucosaminyltransferase led to reduced expression of TF. In contrast, increased cell surface expression of TF was found after ST6GalNAc-II transfection. Thus, expression of TF on the cell surface of SW480 colorectal carcinoma cells depends on the ratio of core 2 .beta.1,6-Nacetylglucosaminyltransferase and ST6GalNAc-II. Earlier immunohistol. studies demonstrated that TF is a prognostic factor for patient survival. Our results suggest that sialyltransferase ST6GalNAc-II is of crucial relevance for the prognostic significance of TF.

REFERENCE COUNT:

THERE ARE 23 CITED REFERENCES AVAILABLE FOR THIS 23 RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 13 OF 43 MEDLINE on STN DUPLICATE 7

ACCESSION NUMBER:

2001189694

DOCUMENT NUMBER:

MEDLINE 21175004 PubMed ID: 11280791

TITLE:

Clinicopathological significance of core 2

beta1,6-N-acetylglucosaminyltransferase messenger RNA

expressed in the pulmonary adenocarcinoma determined by in situ hybridization.

AUTHOR:

Machida E; Nakayama J; Amano J; Fukuda M

CORPORATE SOURCE:

Second Department of Surgery, Shinshu University School of

Medicine, Matsumoto, Japan.

CONTRACT NUMBER:

CA48737 (NCI)

SOURCE:

CANCER RESEARCH, (2001 Mar 1) 61 (5) 2226-31.

Journal code: 2984705R. ISSN: 0008-5472.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

CA33000 (NCI)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

200104

ENTRY DATE:

Entered STN: 20010425

Last Updated on STN: 20010425

Entered Medline: 20010419

AΒ Cell surface carbohydrates of epithelial cells play important roles in tumor progression. Previously, we have shown that expression of core 2 branched O-glycans in colorectal cancer is closely correlated with the vessel invasion and depth of invasion (K. Shimodaira et al., Cancer Res., 57: 5201-5206, 1997). To test whether this is also the case in human lung cancer, we have examined the expression pattern of core 2 beta1,6-N-acetylglucosaminyltransferase (C2GnT) mRNA responsible for the biosynthesis of core 2 branched O-qlycans in 41 cases of lung cancer. Using in situ hybridization, C2GnT mRNA was detected in 73.2% of the lung cancer cells, irrespective of the histopathological type; whereas in normal lung tissues, its expression was restricted to the basal cells of bronchial mucosa. These results indicate that the expression level of C2GnT mRNA was significantly enhanced in association with malignant transformation. Statistical analysis between the C2GnT mRNA expressed in pulmonary adenocarcinoma and clinicopathological variables revealed that the expression of C2GnT was correlated with vessel invasion and lymph node metastasis with significant difference (P < 0.05), but expression of sialyl Le(x), which is frequently expressed in the adenocarcinoma, was not significantly correlated with lymph node metastasis. These results indicate that C2GnT mRNA detected by in situ hybridization reflects the malignant potentials of pulmonary adenocarcinoma, because lymph node metastasis is the most affecting factor to the patients' prognosis.

L12 ANSWER 14 OF 43 BIOSIS COPYRIGHT 2004 BIOLOGICAL ABSTRACTS INC. on STN

ACCESSION NUMBER: DOCUMENT NUMBER:

2002:220540 BIOSIS PREV200200220540

TITLE:

The VNTR polymorphism of human P-selectin

glycoprotein ligand 1 (PSGL-1) affects the kinetics of the PSGL-1-P-selectin bond and the rolling velocity of PSGL-1-

expressing cells in a flow field.

AUTHOR (S):

Afshar-Kharghan, Vahid [Reprint author]; Padilla, Arnoldo [Reprint author]; Romo, Gabriel [Reprint author]; Li, Chester Q. [Reprint author]; Lopez, Jose A. [Reprint

author]

CORPORATE SOURCE:

Thrombosis Research Section, Baylor College of Medicine,

Houston, TX, USA

SOURCE:

Blood, (November 16, 2001) Vol. 98, No. 11 Part 1, pp.

699a. print.

Meeting Info.: 43rd Annual Meeting of the American Society of Hematology, Part 1. Orlando, Florida, USA. December

07-11, 2001. American Society of Hematology.

CODEN: BLOOAW. ISSN: 0006-4971.

DOCUMENT TYPE:

Conference; (Meeting)

Conference; Abstract; (Meeting Abstract)

Conference; (Meeting Poster)

LANGUAGE:

English

ENTRY DATE:

Entered STN: 3 Apr 2002

Last Updated on STN: 3 Apr 2002

Leukocyte adhesion to the vessel wall is an important first step in inflammation. To exit the blood stream, leukocytes first attach and roll on surfaces of activated endothelium or activated platelets. surfaces bear P-selectin, exposed there from internal granule stores, which mediates leukocyte attachment by interacting with its leukocyte counter-receptor, P-selectin glycoprotein ligand-1 (PSGL-1). PSGL-1 is a disulfide-linked homodimer of an elongated mucin-like transmembrane polypeptide that requires posttranslational tyrosine sulfation and carbohydrate core-2 branching and alpha1,3 fucosylation to bind P-selectin. We have recently identified three allelic variants of PSGL-1 due to variable numbers of tandem repeats (VNTR) of a 10-amino acid sequence within the mucin-like core of the molecule. From largest to smallest, the 3 variants are designated A, B, and C, with 16, 15, and 14 repeats, respectively. Of the 16 repeats in variant A, B lacks repeat 2 and C lacks repeats 9 and 10. This polymorphism is expected to affect the length of the polypeptide, and to change the spatial relationship between individual polypeptides in the PSGL-1 dimer if two different variants are coexpressed. To study the role of the VNTR polymorphism in PSGL-1 function, we created Chinese hamster ovary (CHO) cell lines expressing the individual variants, alone and in combinations of two, together with the necessary carbohydrate modifying enzymes, fucosyltransferase VII (FTVII) and the core-2 branching enzyme (C2GnT). We verified by flow cytometry that the individual cell lines express equivalent quantities of surface PSGL-1. We compared cell lines expressing the two forms with the greatest size discrepancy, A and C, for their ability to adhere to and roll on surface-immobilized P-selectin in a parallel-plate flow chamber system, at 2 and 10 dynes/cm2. CHO cells expressing only FTVII and C2GnT served as negative controls. At 2 dynes/cm2, the three PSGL-1-expressing cell lines (CHO AA, CHO CC, and CHO AC) all attached to the P-selectin matrix, with CHO CC rolling 20% faster than CHO AA and 70% faster than CHO AC (p=0.07 and p=0.0002, respectively). At 10 dynes/cm2, CHO CC cells completely failed to attach and roll on the matrix, and CHO AA cells rolled on average 40% faster than CHO AC cells (p=0.004). Thus, the shortest PSGL-1 variant supported cell adhesion more poorly than the longest variant or a combination of the longest and shortest variants, which was the most efficient in mediating adhesion. These data provide evidence that the PSGL-1 VNTR polymorphism may be a significant determinant of leukocyte adhesion in vivo, and thus may also

be a marker for risk of inflammatory disease.

L12 ANSWER 15 OF 43 MEDLINE on STN DUPLICATE 8

ACCESSION NUMBER: 2001503438 MEDLINE

DOCUMENT NUMBER: 21437016 PubMed ID: 11552947

TITLE: Core 2 beta1,6-N-acetylglucosaminyltransferases and

alpha1,3-fucosyltransferases regulate the synthesis of O-glycans on selectin ligands on oral cavity carcinoma

cells.

AUTHOR: Renkonen J; Rabina J; Mattila P; Grenman R; Renkonen R

CORPORATE SOURCE: Department of Bacteriology and Immunology, Haartman

Institute, Haartmaninkatu 3, FIN-0014 University of

Helsinki, Helsinki, Finland.. Jutta.Renkonen@Helsinki.Fi

SOURCE: APMIS, (2001 Jul-Aug) 109 (7-8) 500-6.

Journal code: 8803400. ISSN: 0903-4641.

PUB. COUNTRY: Denmark

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200110

ENTRY DATE: Entered STN: 20010913

Last Updated on STN: 20011015

Entered Medline: 20011011

Selectin-dependent cell binding has importance in the extravasation of AB blood-circulating tumor cells and in the generation of metastases. Cell surface glycoproteins decorated with sialylated, fucosylated epitopes, such as sialyl Lewis(x) (sLe(x)), are ligands for selectins. Not only terminal sLe(x) moieties but also proximal core structures contribute to the formation of binding epitopes for selectins. Core 2 beta1,6-N-acetylglucosaminyltransferases (C2GnT) and alphal, 3-fucosyltransferases (alphal, 3-FucT) have been suggested to be the rate-limiting enzymes in the synthesis of selectin ligands. We analyzed oral cavity epithelial carcinoma cell lines and showed their expression of RNA transcripts for C2GnT and alpha1,3-FucT, identified alpha1,3-FucT enzyme activities, and analyzed the cell surface ${\tt sLe}\left(x\right)$ expression levels. Neither the pattern of expressed enzymes nor the alpha1,3-FucT activity directly predicted the binding capacity of E-selectin. However, only the sLe(x)expressing cell lines were capable of binding to E-selectin, but not to P-selectin, thus putatively promoting the selectin-mediated metastasis. These findings suggest that C2GnT in combination with alpha1,3-Fuc-T contribute to the selectin-mediated metastasis in oral cavity carcinomas.

L12 ANSWER 16 OF 43 BIOSIS COPYRIGHT 2004 BIOLOGICAL ABSTRACTS INC. on STN

ACCESSION NUMBER: 2001:294581 BIOSIS DOCUMENT NUMBER: PREV200100294581

TITLE: Decreased levels of the gel-forming mucin MUC5AC in tears

of Sjogren's Syndrome patients.

AUTHOR(S): Argueso, P. [Reprint author]; Balaram, M. [Reprint author];

Spurr-Michaud, S. [Reprint author]; Dana, R. [Reprint

author]; Gipson, I. K. [Reprint author]

CORPORATE SOURCE: Schepens Eye Research Inst, Harvard Medical School, Boston,

MA, USA

SOURCE: IOVS, (March 15, 2001) Vol. 42, No. 4, pp. S488. print.

Meeting Info.: Annual Meeting of the Association for Research in Vision and Ophthalmology. Fort Lauderdale,

Florida, USA. April 29-May 04, 2001.

DOCUMENT TYPE: Conference; (Meeting)

Conference; Abstract; (Meeting Abstract)

LANGUAGE: English

ENTRY DATE: Entered STN: 20 Jun 2001

Last Updated on STN: 19 Feb 2002

L12 ANSWER 17 OF 43 MEDLINE on STN

ACCESSION NUMBER: 2003471431 MEDLINE

DOCUMENT NUMBER:

PubMed ID: 14533804

TITLE:

Biosynthesis and function of beta 1,6 branched mucin-type

glycans.

AUTHOR:

Beum P V; Cheng P W

CORPORATE SOURCE:

Department of Biochemistry and Molecular Biology,

University of Nebraska Medical Center Omaha, NE 68198-4525,

USA.

CONTRACT NUMBER:

R01 HL48282 (NHLBI)

SOURCE:

Advances in experimental medicine and biology, (2001) 491

279-312. Ref: 167

Journal code: 0121103. ISSN: 0065-2598.

PUB. COUNTRY:

United States

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

General Review; (REVIEW)

(REVIEW, ACADEMIC)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

200311

ENTRY DATE:

Entered STN: 20031010

Last Updated on STN: 20031219 Entered Medline: 20031119

AB The contribution of carbohydrate structure to biomolecular, cellular, and organismal function is well-established, but has not yet received the attention it deserves, perhaps due to the complexity of the structures involved and to a lack of simple experimental methods for relating structure and function. In particular, beta1,6 GlcNAc branching plays a key functional role in processes ranging from inflammation and immune system function to tumor cell metastasis. For instance, synthesis of the core 2 beta1,6 branched structure in the mucin glycan chain by C2GnT enables the expression of functional structures at the termini of polylactosamine chains, such as blood group antigens and sialyl Lewis x. Also, IGnT can create multiple branches on the polylactosamine chain, which may serve as a mechanism for amplifying the functional potency of cell surface glycoproteins and glycolipids. The family of enzymes which creates beta1,6 branched structure in mucin glycans is proving to be quite complex, since multiple isoforms appear to exist for these enzymes, and some of the enzymes are adept at forming more than one type of beta1,6 branched structure, as in the case of C2GnT-M. Furthermore, the enzymes do not appear to be restricted to acting on mucin-type acceptor structures, but are able to act on glycolipid structures as well. Much remains to be learned regarding the specific biological niche filled by each of these enzymes and how their activities complement one another, as well as the manner in which the activities of these enzymes are regulated in the cell.

L12 ANSWER 18 OF 43 SCISEARCH COPYRIGHT 2004 THOMSON ISI on STN

ACCESSION NUMBER: 2001:492533 SCISEARCH

THE GENUINE ARTICLE: 440RF

TITLE:

Regulation of poly-N-acetyllactosamine biosynthesis in

0-glycans

AUTHOR: \

Ujita M (Reprint); Fukuda M

CORPORATE SOURCE:

Burnham Inst, Canc Res Ctr, Glycobiol Program, La Jolla, CA 92037 USA; Meijo Univ, Fac Agr, Dept Appl Biol Chem, Biol Chem Lab, Tempaku Ku, Nagoya, Aichi 4688502, Japan; Meijo Univ, Agr High Tech Res Ctr, Tempaku Ku, Nagoya,

Aichi 4688502, Japan

COUNTRY OF AUTHOR:

USA; Japan

SOURCE:

TRENDS IN GLYCOSCIENCE AND GLYCOTECHNOLOGY, (MAR 2001)

Vol. 13, No. 70, pp. 177-191.

Publisher: FCCA-FORUM CARBOHYDRATES COMING AGE, C/O GAKUSHIN PUBLISHING CO LTD 1-1-8 TARUMI-CHO, SUITA

564-0062, OSAKA, 30015, JAPAN.

ISSN: 0915-7352.

DOCUMENT TYPE:

General Review; Journal

LANGUAGE:

English

REFERENCE COUNT:

73

ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS

Poly-N-acetyllactosamine is a unique carbohydrate composed of N-acetyllactosamine (LacNAc) repeats and provides the backbone structure for additional modifications such as sially Lewis (\mathbf{x}) . It is attached to N-glycans, O-glycans, and glycolipids and synthesized by the alternate addition of beta1,3-linked N-acetylglucosamine (GlcNAc) and beta1,3-linked galactose (Gal) by i-beta1,3-N-acetylglucosaminyltransferase (iGnT) and a member of the beta1,4-galactosyltransferase (beta 4Gal-T) gene family. Poly-N-acetyllactosamines in mucin-type O-glycans can be formed in core 2and core 4-branched oligosaccharides, which are synthesized by core 2 beta1,6-N-acetylglucosaminyltransferase (C2GnT) and core 4 betal,6-N-acetylglucosaminyltransferase (C4GnT), respectively.

beta 4Gal-TIV was found to be most efficient in the addition of a single Gal residue to core 2-branched oligosaccharides among the members of the beta 4Gal-T gene family and to synthesize poly-N-acetyllactosamine in core 2-branched O-glycans together with iGnT. On the other hand, beta 4Gal-TI was shown to be most efficient for poly-N-acetyllactosamine synthesis in N-glycans. In contrast to beta 4Gal-TI, the efficiency of beta 4Gal-TIV decreases dramatically as the accepters contain more LacNAc repeats, consistent with the fact that core 2-branched 0-qlycans contain shorter poly-N-acetyllactosamines than N-glycans in many cells. Poly-N-acetyllactosamines in core 4-branched O-glycans were found to be synthesized most efficiently by iGnT and beta 4Gal-TI although the synthesis in core 4 branches is less efficient than in core 2 branches because of inefficient addition of GlcNAc to core 4 branches by iGnT. Thus, poly-N-acetyllactosamine extension in core 2- and core 4-branched O-glycans is differentially controlled by iGnT and different members of the beta 4Gal-T gene family.

L12 ANSWER 19 OF 43 SCISEARCH COPYRIGHT 2004 THOMSON ISI on STN

ACCESSION NUMBER:

2001:94157 SCISEARCH

THE GENUINE ARTICLE: 395TF

TITLE:

Lipopolysaccharide induces mucus cell metaplasia in mouse

AUTHOR:

Yanagihara K; Seki M; Cheng P W (Reprint)

CORPORATE SOURCE:

Univ Nebraska, Nebraska Med Ctr 984525, Dept Biochem & Mol Biol, 600 S 42nd St, Omaha, NE 68198 USA (Reprint); Univ Nebraska, Nebraska Med Ctr 984525, Dept Biochem & Mol Biol, Omaha, NE 68198 USA; Univ Nebraska, Eppley Canc Ctr, Med Ctr, Omaha, NE 68198 USA

USA

COUNTRY OF AUTHOR:

SOURCE:

AMERICAN JOURNAL OF RESPIRATORY CELL AND MOLECULAR BIOLOGY

(JAN 2001) Vol. 24, No. 1, pp. 66-73.

Publisher: AMER THORACIC SOC, 1740 BROADWAY, NEW YORK, NY

10019-4374 USA. ISSN: 1044-1549.

DOCUMENT TYPE:

Article; Journal

LANGUAGE:

English

REFERENCE COUNT:

ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS

ABA murine model of lipopolysaccharide (LPS)-induced airway inflammation and epithelial cell phenotypic change, and the time courses of these events are described. A single intratracheal instillation of Pseudomonas aeruginoso LPS in mice resulted in massive recruitment of neutrophils to the lung 2 d after treatment as assessed by differential cell counts of the inflammatory cells in bronchoalveolar lavage fluid and histologic assessment of hemotoxylin and eosin (H&E)-stained lung sections. The LPS-induced neutrophilic inflammation subsided substantially on Day 4 and essentially vanished by Day 7. Airway epithelial mucus cells were not detected by Alcian blue periodic acid-Schiff staining until Day 4 after

LPS treatment and became more abundant in number as well as in mucus content on Day 7. The expression of Muc5ac messenger RNA (mRNA) as well as glycoprotein was enhanced on Day 2, peaked on Day 4 and decreased on Day 7, whereas enhanced expression of mucin core 2 beta6 N-acetylglucosaminyltransferase (C2GnT)-M mRNA was not detected until Day 4 and peaked on Day 7. The expression of . C2CnT-L mRNA in the lung, a marker for activated leukocytes as well as mucus cells, peaked on Day 2 and remained moderately high until Day 7. C2GnT-L mRNA expression in LPS-treated lung correlated with the presence of neutrophils and the appearance of mucus cells in the airway epithelium. We conclude that mucus cell metaplasia and hyperplasia can be generated in mouse lungs with a single intratracheal instillation of LPS. In addition, C2GnT-M may serve as a marker for mucus cells in mouse lung. This LPS-induced mucus cell metaplasia and hyperplasia model should be useful for the study of Pseudomonos-induced airway mucus hypersecretory diseases.

L12 ANSWER 20 OF 43 HCAPLUS COPYRIGHT 2004 ACS on STN

ACCESSION NUMBER:

2000:754450 HCAPLUS

DOCUMENT NUMBER:

133:319062

TITLE:

Cloning and characterization of human and mouse .beta.-1-6-N-

acetylglucosaminyltransferase that forms core 2, core

4 and I branches and its functional fragments

INVENTOR(S):

Fukuda, Minoru; Yeh, Jiunn-Chern

PATENT ASSIGNEE(S):

The Burnham Institute, USA

SOURCE:

U.S., 30 pp. CODEN: USXXAM

DOCUMENT TYPE:

Patent

LANGUAGE:

English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE	
	US 6136580 PRITY APPLN. INFO.	:		US 1999-233506	19990119	
AB	.beta1.fwdarw. I branching actiinvention provide nucleic acids, a and methods involved an acceptor mol.	6-N-activities les sub intibod olving by co	etylglucosam s, designated estantially p lies specific such compns. entacting the	ates to a novel, multi-functional lglucosaminyltransferase with core 2, core 4 and esignated C2GnT-M. In particular, the ntially pure C2GnT-M polypeptides and specifically reactive with C2GnT-M, h compns. Also provided is a method of modifying cting the acceptor mol. with a substantially pure		
	C2GnT-M polypeptide or a functional fragment under conditions that allow addn. of core 2, core 4 or I GlcNAc linkages to the acceptor and an acceptor mol. produced by the method. A cDNA and encoded amacid sequence of human C2GnT-M are disclosed.					
	50,963 Da and a provides function	38 ami predic	no acid resideted type II agments and	cterized as a dues having a predict membrane topol. The derivs. of C2GnT -M th s of full-length C2G n	invention also hat have	
,	biol. activity of any combination acetylglucosamin	of a C2 of the nyltran	GnT-M fragme core 2 .bet sferase, cor	nt can be, for examp a1.fwdarw.6-N- e 4 .beta1.fwdarw.6 -branching .beta1.:	le, one or 6-N-	
	acetylglucosaminyltransferase activities of C2GnT-M. The invention also provides a substantially pure nucleic acid mol. encoding C2GnT-M or a functional fragment or deriv. thereof, or the complement of the nucleic acid mol., wherein C2GnT-M is					
	characterized as .beta1.fwdarw.	a pol 6-N-ac	ypeptide hav. etylglucosam	ing core 2, core 4 and involtransferase actives contg. nucleic acid	nd I branching vities. Also	

C2GnT-M or a functional fragment or deriv. thereof. The cDNA and encoded amino acid sequences of the mouse C2GnT-M CCL1 and IS3 fragments are provided. The invention further provides an antibody or antigen binding fragment thereof that is specifically reactive with

C2GnT-M or with a functional fragment or deriv. hereof.

REFERENCE COUNT: 21 THERE ARE 21 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L12 ANSWER 21 OF 43 MEDLINE on STN DUPLICATE 9

ACCESSION NUMBER: 2000283898 MEDLINE

DOCUMENT NUMBER: 20283898 PubMed ID: 10811884

TITLE: A multipotential beta -1,6-N-acetylqlucosaminyl-transferase

is encoded by bovine herpesvirus type 4.

AUTHOR: Vanderplasschen A; Markine-Goriaynoff N; Lomonte P; Suzuki

M; Hiraoka N; Yeh J C; Bureau F; Willems L; Thiry E; Fukuda

M; Pastoret P P

CORPORATE SOURCE: Department of Immunology-Vaccinology (B43 bis), Faculty of

Veterinary Medicine, University of Liege, B-4000 Liege,

Belgium.. A.vdplasschen@ulg.ac.be

CONTRACT NUMBER: R37 CA33000 (NCI)

SOURCE: PROCEEDINGS OF THE NATIONAL ACADEMY OF SCIENCES OF THE

UNITED STATES OF AMERICA, (2000 May 23) 97 (11) 5756-61.

Journal code: 7505876. ISSN: 0027-8424.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals; AIDS

OTHER SOURCE: GENBANK-AF231105

ENTRY MONTH: 200006

ENTRY DATE: Entered STN: 20000714

Last Updated on STN: 20000714 Entered Medline: 20000630

AB The beta-1,6-N-acetylglucosaminyltransferase (beta1,6GnT) gene family encodes enzymes playing crucial roles in glycan synthesis. Important changes in beta1,6GnT expression are observed during development, oncogenesis, and immunodeficiency. The most characterized beta1,6GnTs in this gene family are the human (h) C2GnT -L and h-IGnT, which have core 2 [Galbeta1-->3 (GlcNAcbeta1-->6) GalNAc] and I branching [GlcNAcbeta1-->3(GlcNAcbeta1-->6)Gal] activities, respectively. Recently, h-C2GnT-M was shown to be unique in forming core 2, core 4 [GlcNAcbeta1-->3(GlcNAcbeta1-->6)GalNAc], and I structures. To date, the beta1,6GnT gene family has been characterized only in mammals. Here, we describe that bovine herpesvirus type 4 (BHV-4) encodes a betal, 6GnT expressed during viral replication and exhibiting all of the core 2, core 4, and I branching activities. Sequencing of the BHV-4 genome revealed an ORF, hereafter called BORFF3-4, encoding a protein (pBORFF3-4) exhibiting 81.1%, 50.7%, and 36.6% amino acid identity with h-C2GnT-M, h-C2GnT-L, and h-IGnT, respectively. Reverse transcriptase-PCR analysis revealed that BORFF3-4 is expressed during BHV-4 replication. Expression of BORFF3-4 in Chinese hamster ovary cells directed the expression of core 2 branched oligosaccharides and I antigenic structures on the cell surface. Moreover, a soluble form of pBORFF3-4 had core 4 branching activity in addition to core 2 and I branching activities. Finally, infection of a C2GnT-negative cell line with BHV-4 induced expression of core 2 branched oligosaccharides. This study extends the betal,6GnT gene family to a viral gene and provides a model to study the biological functions of a beta1,6GnT in the context of viral infection.

L12 ANSWER 22 OF 43 MEDLINE on STN DUPLICATE 10

ACCESSION NUMBER: 2000161141 MEDLINE

DOCUMENT NUMBER: 20161141 PubMed ID: 10694812

TITLE: Expression of human Wiskott-Aldrich

syndrome protein in patients' cells leads to partial correction of a phenotypic abnormality of cell surface

glycoproteins.

AUTHOR: Huang M M; Tsuboi S; Wong A; Yu X J; Oh-Eda M; Derry J M;

Francke U; Fukuda M; Weinberg K I; Kohn D B

Division of Research Immunology/Bone Marrow CORPORATE SOURCE:

Transplantation, Childrens Hospital Los Angeles, Los

Angeles, CA 90033, USA.

CONTRACT NUMBER: DK09430-02 (NIDDK)

> R01-DK49000 (NIDDK) R37-CA33000 (NCI)

SOURCE: GENE THERAPY, (2000 Feb) 7 (4) 314-20.

Journal code: 9421525. ISSN: 0969-7128.

PUB. COUNTRY: ENGLAND: United Kingdom

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals; AIDS

ENTRY MONTH: 200003

ENTRY DATE: Entered STN: 20000327

> Last Updated on STN: 20000327 Entered Medline: 20000316

AB The Wiskott-Aldrich syndrome (WAS) is an uncommon X-linked recessive disease characterized by thrombocytopenia, eczema and immunodeficiency. The biochemical defect of this disorder primarily affects cells derived from bone marrow. To understand better the molecular mechanisms underlying this disease and to evaluate the possibility of correcting the genetic defects in hematopoietic cells, a Moloney murine leukemia virus (MoMLV) - based retroviral vector carrying a functional Wiskott-Aldrich syndrome protein (WASp) cDNA driven by an SV40 promoter (LNS-WASp) was constructed. A packaging cell line containing this vector produced a stable level of WAS protein and maintained a high titer of viral output. Epstein-Barr virus (EBV)-transformed B lymphoblastoid cell lines (B-LCL) from WAS patients, which lack expression of the WAS protein, were transduced by the LNS-WASp retroviral vector and showed expression of WASp by Western blot. Analysis of the O-glycan pattern on cell surface glycoproteins from WAS patients' B-LCL showed an altered glycosylation pattern, due to increased activity of beta-1, 6-N-acetylglucosaminyltransferase (C2GnT). Transduction by the retroviral vector carrying the functional WASp cDNA partially restored the abnormal glycosylation pattern, and was accompanied by a decreasing C2GnT activity. These findings imply a functional linkage between the WAS protein and the expression of the glycosyltransferase involved in the O-glycosylation, and also suggest a potential gene therapy via transferring a functional WASp cDNA into hematopoietic cells for Wiskott-Aldrich syndrome. Gene Therapy (2000) 7, 314-320.

L12 ANSWER 23 OF 43 MEDLINE on STN DUPLICATE 11

ACCESSION NUMBER: 2000211217 MEDLINE

DOCUMENT NUMBER: 20211217 PubMed ID: 10745191

TITLE: Engineering of coordinated up- and down-regulation of two

glycosyltransferases of the O-glycosylation pathway in

Chinese hamster ovary (CHO) cells.

AUTHOR: Prati E G; Matasci M; Suter T B; Dinter A; Sburlati A R;

Bailey J E

CORPORATE SOURCE: Institute of Biotechnology, ETH Zurich, CH-8093 Zurich,

SOURCE: BIOTECHNOLOGY AND BIOENGINEERING, (2000 May 5) 68 (3)

Journal code: 7502021. ISSN: 0006-3592.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200005 ENTRY DATE:

Entered STN: 20000613

Last Updated on STN: 20000613 Entered Medline: 20000531

AΒ Production of O-linked oligosaccharides that interact with selectins to mediate cell-cell adhesion occurs in one segment of a branched glycan biosynthesis network. Prior efforts to direct the branched pathway towards selectin-binding oligosaccharides by amplifying enzymes in this branch of the network have had limited success, suggesting that metabolic engineering to simultaneously inhibit the competing pathway may also be required. We report here the partial cloning of the CMP-sialic acid:Galbeta1,3GalNAcalpha2, 3-sialyltransferase (ST3Gal I) gene from Chinese hamster ovary (CHO) cells and the simultaneous inhibition of expression of CHO cell ST3Gal I gene and overexpression of the human UDP-GlcNAc:Galbeta1, 3GalNAc-R beta1,6-Nacetylglucosaminyltransferase (C2GnT) gene. A tetracycline-regulated system adjoined to tricistronic expression technology allowed "one-step" transient manipulation of multiple enzyme activities in the O-glycosylation pathway of a previously established CHO cell line already engineered to express alpha1, 3-fucosyltransferase VI (alpha1,3-Fuc-TVI). Tetracycline-regulated coexpression of a ST3Gal I fragment, cloned in the antisense orientation, and of C2GnT cDNA resulted in inhibition of the ST3Gal I enzymatic activity and increase in C2GnT activity which varied depending on the extent of tetracycline reduction in the cell culture medium. This simultaneous regulated inhibition and activation of the two key enzyme activities in the O-glycosylation pathway of mammalian cells is an important addition to the metabolic engineering Copyright 2000 John Wiley & Sons, Inc.

ACCESSION NUMBER:

L12 ANSWER 24 OF 43 BIOSIS COPYRIGHT 2004 BIOLOGICAL ABSTRACTS INC. on STN 2001:311327 BIOSIS

DOCUMENT NUMBER:

PREV200100311327

TITLE:

Regulation of cell surface sialyl-LeX expression

level in human B cell precursor leukemia.

AUTHOR (S):

Nakamura, Mitsuru [Reprint author]; Furukawa, Yusuke

[Reprint author]; Matsuda, Michio [Reprint author]

CORPORATE SOURCE:

Cell and Molecular Medicine, Jichi Medical School,

Minamikawachi, Japan

SOURCE:

Blood, (November 16, 2000) Vol. 96, No. 11 Part 2, pp. 43b.

print.

Meeting Info.: 42nd Annual Meeting of the American Society of Hematology. San Francisco, California, USA. December 01-05, 2000. American Society of Hematology.

CODEN: BLOOAW. ISSN: 0006-4971.

DOCUMENT TYPE:

Conference; (Meeting)

Conference; Abstract; (Meeting Abstract)

LANGUAGE:

English

ENTRY DATE:

Entered STN: 27 Jun 2001

Last Updated on STN: 19 Feb 2002

AB Cell surface sialyl-LeX (sLeX) is known as one of established selectin ligands and biosynthesized through the action of alpha1->3 fucosyltransferase VII (FucTVII) and alpha2->3sialyltransferase IV (ST3GalIV). Significant down regulation of sLeX antigen expression during pre-B cell differentiation is mediated by leukocyte type core 2 beta1->6GlcNAc-transferase (C2GnT or C2GnT/L). We have examined further role of C2GnT/L comparing with FucTVII, ST3GalIV, and mucin type of C2GnT (C2GnT/M) by establishing gene overexpressed sublines from pre-B lymphoid KM3 cells. While sLeX down regulation was partially blocked in the transfectants of FucTVII, ST3GalIV, and C2GnT/M, the block was far from sufficient. Only C2GnT/L could completely block the suppression of sLeX expression. Inability of C2GnT /M for substituting C2GnT/L was further confirmed using Tet-ON

gene expression system. For clinical samples, lymphoblasts from B-ALL patients exhibited high sLeX expression and strong message expression of FucTVII, ST3GalIV, and C2GnT/L. By contrast in B-CLL lymphoblasts, sLeX was not expressed and all glycosyltransferase messages were suppressed as well as C2GnT/L. These suggest that, before the suppression of FucTVII and ST3GalIV, the key C2GnT/L is first down regulated during human B cell differentiation resulting sufficient decrease of sLeX antigen expression. Furthermore, it is suggested that this role of C2GnT is solely played by leukocyte type C2GnT/L and cannot be replaced by another type of the enzyme, C2GnT/M.

ANSWER 25 OF 43 BIOTECHDS COPYRIGHT 2004 THOMSON DERWENT/ISI on STN

ACCESSION NUMBER: 2000-02839 BIOTECHDS

TITLE: Glycosylation engineering in Chinese hamster ovary cells

using tricistronic vectors;

expression of core 2 N-

acetylglucosaminyltransferase and alpha-1,3-

fucosyltransferase-III or -IV in CHO cell culture for

production of glycoprotein having a human-like

glycosylation pattern

Dinter A; Zeng S; Berger B; *Berger E G

CORPORATE SOURCE: Univ.Zurich-Inst.Physiol.

LOCATION:

Institute of Physiology, University of Zurich, Winterthurerstrasse 190, CH-8057 Zurich, Switzerland.

Email: egberger@physiol.unizh.ch

Biotechnol.Lett.; (2000) 22, 1, 25-30 SOURCE:

> CODEN: BILED3 ISSN: 0141-5492

DOCUMENT TYPE: Journal

LANGUAGE: English

CHO cells lack core-2 N-acetylglucosaminyltransferase (C2GnT) AB and alpha-1,3-fucosyltransferase (a3FucT), 2 key enzymes for the expression of O-linked sialyl-Lewis-X structures which are required for P-selectin binding. Tricistronic vectors were constructed that encoded the neomycin-resistance marker in the 3rd cistron, and mouse C2GnT or a3FucT-III or -IV in either the 1st or 2nd cistron, respectively, under control of the SV40 virus promoter. The separate moieties of the constructs were separated by internal ribosomal entry site (IRES) sequences. CHO cells transfected with the tricistronic vectors produced C2GnT and a3FucT activities in a constant ratio. P-selectin binding of recombinant P-selectin glycoprotein ligand-1 (PGSL-1) was demonstrated in CHO cells stably expressing a3FucT-IRES-C2GnT or a3FucT-III-IRES-C2GnT and cotransfected with a soluble form of PGSL-1 and a puromycin-resistance conferring plasmid. This approach appears to be well suited to engineering glycosylation pathways that require matched transfer rates of sequentially active glycosyltransferase, e.g. for production of recombinant glycoproteins having human -like glycosylation patterns. (10 ref)

MEDLINE on STN L12 ANSWER 26 OF 43 DUPLICATE 12

ACCESSION NUMBER:

1999386938 MEDLINE

DOCUMENT NUMBER: 99386938

PubMed ID: 10455130

TITLE:

Expression of core 2 beta-1,6-N-

acetylqlucosaminyltransferase in a human pancreatic cancer cell line results in altered expression of MUC1 tumor-associated epitopes.

AUTHOR: CORPORATE SOURCE: Beum P V; Singh J; Burdick M; Hollingsworth M A; Cheng P W

Department of Biochemistry and Molecular Biology,

University of Nebraska Medical Center, Omaha, Nebraska

68198, USA.

CONTRACT NUMBER: HL48242 (NHLBI)

P30 CA36727 (NCI)

RO1 CA69234 (NCI)

SOURCE: JOURNAL OF BIOLOGICAL CHEMISTRY, (1999 Aug 27) 274 (35)

24641-8.

Journal code: 2985121R. ISSN: 0021-9258.

PUB. COUNTRY:

United States

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

199909

ENTRY DATE:

Entered STN: 19991012

Last Updated on STN: 19991012 Entered Medline: 19990930

AB Many tumor-associated epitopes possess carbohydrate as a key component, and thus changes in the activity of glycosyltransferases could play a role in generating these epitopes. In this report we describe the stable transfection of a human pancreatic adenocarcinoma cell line, Panc1-MUC1, with the cDNA for mucin core 2 GlcNAc-transferase (C2GnT), which creates the core 2 beta-1,6 branch in mucin-type glycans. These cells lack endogenous C2GnT activity but

express a recombinant human MUC1 cDNA.

C2GnT-transfected clones expressing different

levels of C2GnT were characterized using monoclonal antibodies CC49, CSLEX-1, and SM-3, which recognize tumor-associated epitopes.

Increased C2GnT expression led to greatly diminished expression of the CC49 epitope, which we identified as

NeuAcalpha2,6(Galbeta1,3)GalNAcalpha-Ser/Thr in the Panc1-MUC1 cells.

This was accompanied by the emergence of the CSLEX-1 epitope, sialyl Lewis x (NeuAcalpha2,3Galbeta1,4(Fucalpha1,3)GlcNAc-R), an important selectin ligand. Despite this, however, the C2GnT transfectants could

not bind to selectins. Increased C2GnT expression

also led to masking of the SM-3 peptide epitope, which persisted after the removal of sialic acid, further suggesting greater complexity of the core 2-associated O-glycans on MUC1. The results of this study suggest that C2GnT could play a regulatory role in the expression of

certain tumor-associated epitopes.

L12 ANSWER 27 OF 43 MEDLINE on STN **DUPLICATE 13**

ACCESSION NUMBER:

1999143102

MEDLINE

DOCUMENT NUMBER:

99143102 PubMed ID: 9988682

TITLE:

Control of O-glycan branch formation. Molecular

cloning of human cDNA encoding a novel

beta1,6-N-acetylglucosaminyltransferase forming core 2 and

core 4.

AUTHOR:

Schwientek T; Nomoto M; Levery S B; Merkx G; van Kessel A

G; Bennett E P; Hollingsworth M A; Clausen H

CORPORATE SOURCE:

School of Dentistry, University of Copenhagen, Norre Alle

20, 2200 Copenhagen N, Denmark.

CONTRACT NUMBER:

1 RO1 CA66234 (NCI)

1RO1 CA66234 (NCI)

5 P41 RR05351 (NCRR)

SOURCE: JOURNAL OF BIOLOGICAL CHEMISTRY, (1999 Feb 19) 274 (8)

Journal code: 2985121R. ISSN: 0021-9258.

PUB. COUNTRY:

United States

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: OTHER SOURCE:

Priority Journals GENBANK-AF038650

ENTRY MONTH:

199903

ENTRY DATE:

Entered STN: 19990326

Last Updated on STN: 20000303 Entered Medline: 19990318

AΒ A novel human UDP-GlcNAc:Gal/GlcNAcbeta1-3GalNAcalpha beta1,

6GlcNAc-transferase, designated C2/4GnT, was identified by BLAST analysis

of **expressed** sequence tags. The sequence of C2/4GnT encoded a putative type II transmembrane protein with significant sequence similarity to **human C2GnT** and IGnT.

Expression of the secreted form of C2/4GnT in insect cells showed that the gene product had UDP-N-acetyl-alpha-D-glucosamine:acceptor beta1, 6-N-acetylglucosaminyltransferase (beta1,6GlcNAc-transferase) activity. Analysis of substrate specificity revealed that the enzyme catalyzed O-glycan branch formation of the core 2 and core 4 type. NMR analyses of the product formed with core 3-para-nitrophenyl confirmed the product core 4-para-nitrophenyl. The coding region of C2/4GnT was contained in a single exon and located to chromosome 15q21.3. Northern analysis revealed a restricted expression pattern of C2/4GnT mainly in colon, kidney, pancreas, and small intestine. No expression of C2/4GnT was detected in brain, heart, liver, ovary, placenta, spleen, thymus, and peripheral blood leukocytes. The **expression** of core 2 O-glycans has been correlated with cell differentiation processes and cancer. The results confirm the predicted existence of a betal,6GlcNAc-transferase that functions in both core 2 and core 4 0-glycan branch formation. redundancy in beta1,6GlcNAc-transferases capable of forming core 2 O-glycans is important for understanding the mechanisms leading to specific changes in core 2 branching during cell development and malignant transformation.

L12 ANSWER 28 OF 43 MEDLINE on STN DUPLICATE 14

ACCESSION NUMBER:

1999115671 · MEDLINE

DOCUMENT NUMBER:

99115671 PubMed ID: 9915862

TITLE:

Molecular cloning and expression of a

novel beta-1, 6-N-acetylglucosaminyltransferase that forms

core 2, core 4, and I branches.

AUTHOR:

Yeh J C; Ong E; Fukuda M

CORPORATE SOURCE:

Glycobiology Program, Cancer Research Center, the Burnham

Institute, La Jolla, California 92037, USA.

CONTRACT NUMBER:

PO1 CA71932 (NCI)

R37 CA33000 (NCI)

SOURCE:

JOURNAL OF BIOLOGICAL CHEMISTRY, (1999 Jan 29) 274 (5)

3215-21.

Journal code: 2985121R. ISSN: 0021-9258.

PUB. COUNTRY:

DOCUMENT TYPE:

United States
Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English
Priority Journals

FILE SEGMENT: OTHER SOURCE: Priority Journals GENBANK-AF102542

ENTRY MONTH:

199903

ENTRY DATE:

Entered STN: 19990316

Last Updated on STN: 19990316 Entered Medline: 19990303

Mucin-type O-glycans are classified according to their core structures. AB Among them, cores 2 and 4 are important for having N-acetyllactosamine side chains, which can be further modified to express various functional oligosaccharides. Previously, we discovered by cloning cDNAs that the core 2 branching enzyme, termed core 2 beta-1,6-Nacetylglucosaminyltransferase-leukocyte type (C2GnT-L), is highly homologous to the I branching beta-1, 6-Nacetylqlucosaminyltransferase (IGnT) (Bierhuizen, M. F. A., Mattei, M.-G., and Fukuda, M. (1993) Genes Dev. 7, 468-478). Using these homologous sequences as probes, we identified an expressed sequence tag in dbEST, which has significant homology to C2GnT-L and IGnT. This approach, together with 5'and 3' rapid amplification of cDNA ends, yielded a human cDNA that encompasses a whole coding region of an enzyme, termed C2GnT-mucin type (C2GnT -M). C2GnT-M has 48.2 and 33.8% identity with C2GnT-L and IGnT at the amino acid levels. The expression of C2GnT-M cDNA directed the expression of core 2 branched oligosaccharides and I antigen on the cell surface. Moreover, a soluble chimeric C2GnT-M had core 4 branching activity in addition to core 2 and I branching activities. A soluble chimeric C2GnT-L, in contrast, almost exclusively contains core 2 branching activity. Northern blot analysis demonstrated that the C2GnT-M transcripts are heavily expressed in colon, small intestine, trachea, and stomach, where mucin is produced. In contrast, the transcripts of C2GnT-L were more widely detected, including the lymph node and bone marrow. These results indicate that the newly cloned C2GnT-M plays a critical role in O-glycan synthesis in mucins and might have distinctly different roles in oligosaccharide ligand formation compared with C2GnT-L.

L12 ANSWER 29 OF 43 MEDLINE on STN

DUPLICATE 15

ACCESSION NUMBER:

1999166954

MEDLINE

DOCUMENT NUMBER:

99166954 PubMed ID: 10069424

TITLE:

Core 2-containing O-glycans on CD43 are preferentially

expressed in the memory subset of human

CD4 T cells.

AUTHOR:

Mukasa R; Homma T; Ohtsuki T; Hosono O; Souta A; Kitamura

T; Fukuda M; Watanabe S; Morimoto C

CORPORATE SOURCE:

Department of Clinical Immunology and AIDS Research Center,

Institute of Medical Science, University of Tokyo, Japan.

CONTRACT NUMBER: AI29530 (NIAID)

AR33713 (NIAMS) CA33000 (NCI)

SOURCE:

INTERNATIONAL IMMUNOLOGY, (1999 Feb) 11 (2) 259-68.

Journal code: 8916182. ISSN: 0953-8178.

PUB. COUNTRY:

ENGLAND: United Kingdom

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

199908

ENTRY DATE:

Entered STN: 19990827

Last Updated on STN: 19990827

Entered Medline: 19990817

AΒ Human CD4 T cells can be divided into two functionally distinct subsets: a CD45RO+ memory subset and a CD45RA+ naive subset. In an attempt to identify novel cell surface molecules on these cells, we have developed a mAb, anti-1D4. The antigen defined by anti-1D4 was preferentially expressed on the memory subset of freshly isolated peripheral CD4 T cells and 1D4+ CD4 T cells functionally corresponded to memory T cells. Retrovirus-mediated expression cloning revealed that the 1 D4 antigen is human CD43. Transfection of CHO-leu cells, which stably express human CD43, with core 2 beta-1,6-N-acetylglucosaminyltransferase (C2GnT) conferred expression of the 1D4 antigen and mRNA of C2GnT was detected by RT-PCR only in 1D4+ T cells but not in 1D4 - T cells, implying that the 1 D4 antigen is composed of core 2-containing O-glycans on CD43. Reactivity with anti-1 D4 was completely abolished when cells were treated with neuraminidase, while them remained weak binding of anti-T305, a previously described mAb which also reacts with CD43 modified with core 2-containing O-glycans. Moreover, anti-1D4 markedly reacted with NIH-3T3 cells expressing human. CD43 and low levels of endogenous C2GnT, whereas anti-T305 reacted slightly. These results indicate that the 1D4 antigen is distinct from the epitope defined by anti-T305 and anti-1D4 is a more sensitive probe to detect core 2-containing 0-glycans than anti-T305. Taken together, our results indicate that core 2-containing O-glycans, whose expression can easily be detected with anti-1D4, are preferentially expressed in the CD45RO+ memory subset of CD4 T cells.

DOCUMENT NUMBER: 20069459 PubMed ID: 10601651

Simultaneous core 2 beta1-->6N-TITLE:

acetylglucosaminyltransferase up-regulation and sialyl-Le(X) expression during activation of

human tonsillar B lymphocytes.

Nakamura M; Ishida T; Kikuchi J; Furukawa Y; Matsuda M AUTHOR:

Division of Molecular Hemopoiesis, Center for Molecular CORPORATE SOURCE:

Medicine, Jichi Medical School, Minamikawachi, Tochiqi,

Japan.. owlmnaka@jichi.ac.jp

SOURCE: FEBS LETTERS, (1999 Dec 10) 463 (1-2) 125-8.

Journal code: 0155157. ISSN: 0014-5793.

PUB. COUNTRY: Netherlands

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200002

ENTRY DATE: Entered STN: 20000209

Last Updated on STN: 20000209 Entered Medline: 20000203

We have investigated the regulation mechanism of the surface sialy1-Le(X) AB (sLe(X)) expression level in tonsillar B cells during activation. sLe(X) antigen became strongly positive after activation, while resting B cells were weakly positive. sLe(X) structures were mainly located on O-linked oligosaccharide chains of glycoprotein. Transcripts of FucT-VII and core 2 GlcNAc transferase (C2GnT) were up-regulated after activation, while those of ST3GalIV and beta1-->4GalT-I were expressed constitutively. However, the up-regulation of C2GnT was more dramatic than that of FucT-VII. These results suggest that sLe(X) expression level is regulated by

L12 ANSWER 31 OF 43 HCAPLUS COPYRIGHT 2004 ACS on STN

C2GnT during tonsillar B cell activation.

ACCESSION NUMBER: 1999:770783 HCAPLUS

DOCUMENT NUMBER: 132:103285

Effects of TGF-.beta.1 on .beta.1,6N-TITLE:

acetylglucosaminyltransferase activity in the

human colonic cancer cell line Caco-2

AUTHOR(S): Kumano, Koji

CORPORATE SOURCE: Second Department of Internal Medicine, Osaka Medical

College, Japan

SOURCE: Osaka Ika Daigaku Zasshi (1999), 58(2), 10-20

CODEN: OIDZAU; ISSN: 0030-6118

PUBLISHER: Osaka Ika Daigaku Igakkai

DOCUMENT TYPE: Journal

LANGUAGE: Japanese

The effects of TGF-.beta.1 on .beta.1-6 N-acetylglucosaminyltransferase (C2GnT) activity in the human colonic cancer cell line,

Caco-2 (adenocarcinoma), was examd. C2GnT activity was assayed with various concns. (0,1,2,5,10 ng / ml) of TGF-.beta.1. The activity increased in proportion to the concn. of TGF-.beta.1 up to 5 ng / ml and reached a max. at 5 ng/mL. The activity increased gradually after 24 h incubation with 5 ng/mL of TGF-.beta.1 and remained at this level until 72 h. RT-PCR showed strong expression of C2GnT mRNA

after addn. of 5 ng/mL or 10 ng/mL of TGF-.beta.1. This was consistent with the obsd. increase in enzyme activity. Both TGF-.beta. type I and type II receptors were detected by immunofluorescence. TGF-.beta.1 inhibited growth of Caco-2 cells suggesting that transduction of the TGF-/.beta.1 signal functioned properly. In conclusion, TGF-.beta.1

increased C2GnT mRNA levels and enzyme activity. C2GnT activity is controlled at the mRNA level.

L12 ANSWER 32 OF 43 MEDLINE on STN DUPLICATE 17

ACCESSION NUMBER: 1998434594 MEDLINE

DOCUMENT NUMBER: 98434594 PubMed ID: 9756922 TITLE:

Single glycosyltransferase, core 2 beta1-->6-N-

acetylglucosaminyltransferase, regulates cell surface

sialyl-Lex expression level in human

pre-B lymphocytic leukemia cell line KM3 treated with

phorbolester.

AUTHOR:

Nakamura M; Kudo T; Narimatsu H; Furukawa Y; Kikuchi J;

Asakura S; Yang W; Iwase S; Hatake K; Miura Y

CORPORATE SOURCE:

Division of Hemopoiesis, Jichi Medical School,

Minamikawachi, Tochigi 329-04, Japan.

SOURCE:

JOURNAL OF BIOLOGICAL CHEMISTRY, (1998 Oct 9) 273 (41)

26779-89.

Journal code: 2985121R. ISSN: 0021-9258.

PUB. COUNTRY:

United States

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

199811

ENTRY DATE:

Entered STN: 19990106

Last Updated on STN: 19990106 Entered Medline: 19981102

AB Sialyl-Lex (sLex) antigen expression recognized by KM93 monoclonal antibody was significantly down-regulated during

differentiation induced by 12-0-tetradecanoylphorbol-13-acetate (TPA) in

human pre-B lymphocytic leukemia cell line KM3. The sLex determinants were almost exclusively expressed on O-linked

oligosaccharide chains of an O-glycosylated 150-kDa glycoprotein (gp150).

A low shear force cell adhesion assay showed that TPA treatment

significantly inhibited E-selectin-mediated cell adhesion. Transcript

and/or enzyme activity levels of alpha1-->3-fucosyltransferase,

alpha2-->3-sialyltransferase, beta1-->4-galactosyltransferase, and

elongation beta1-->3-N-acetylglucosaminyltransferase did not correlate

with sLex expression levels. However, transcript and enzyme activity levels of core 2 GlcNAc-transferase (C2GnT) were

significantly down-regulated during TPA treatment. Following transfection

and constitutive expression of full-length exogenous

C2GnT transcript, C2GnT enzyme activities were

maintained at high levels even after TPA treatment and down-regulation of

cell surface sLex antigen expression by TPA was completely

abolished. Furthermore, in the transfected cells, the KM93 reactivity of gp150 was not reduced by TPA treatment, and the inhibition of cell

adhesion by TPA was also blocked. These results suggest that sLex expression is critically regulated by a single

glycosyltransferase, C2GnT, during differentiation of KM3 cells.

L12 ANSWER 33 OF 43

MEDLINE on STN

DUPLICATE 18

ACCESSION NUMBER: DOCUMENT NUMBER:

1998250146 MEDLINE 98250146 PubMed ID: 9590264

TITLE:

An sLex-deficient variant of HL60 cells exhibits high levels of adhesion to vascular selectins: further evidence that HECA-452 and CSLEX1 monoclonal antibody epitopes are

not essential for high avidity binding to vascular

AUTHOR:

Wagers A J; Stoolman L M; Craig R; Knibbs R N; Kansas G S Department of Microbiology-Immunology, Northwestern Medical

School, Chicago, IL 60611, USA. AI 33189 (NIAID)

CONTRACT NUMBER:

HL 31963 (NHLBI) SOURCE:

CORPORATE SOURCE:

JOURNAL OF IMMUNOLOGY, (1998 May 15) 160 (10) 5122-9.

Journal code: 2985117R. ISSN: 0022-1767.

PUB. COUNTRY:

United States

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Abridged Index Medicus Journals; Priority Journals

ENTRY MONTH:

199805

ENTRY DATE:

Entered STN: 19980609

Last Updated on STN: 19980609 Entered Medline: 19980528

AΒ Selectins are carbohydrate-binding cell adhesion molecules that play a key role in the initiation of inflammatory responses. Several studies have suggested that the sialylated, fucosylated tetrasaccharide sialyl Lewis X (sLex) is an important component of leukocyte ligands for E- and P-selectin. We have identified a stable variant of the HL60 cell line, HL60var, which displays a nearly complete absence of staining with several mAb directed against sLex and/or sLex-related structures. HL60var also exhibits a concomitant increase in reactivity with mAb directed against the unsialylated Lewis X (Lex/CD15) structure. Despite this sLex deficiency, HL60var binds well to both E- and P-selectin. No significant differences in expression of alpha1,3-fucosyltransferases, C2GnT (Core2 transferase), or P-selectin glycoprotein ligand-1 between HL60var and typical sLex(high) HL60 cells were detected. Although the precise molecular basis for the sLex(-/low) phenotype of HL60var remains uncertain, flow cytometric analysis with the sialic acid-specific Limax flavus lectin revealed a sharp reduction in HL60var surface sialylation. Thus, the loss in mAb reactivity may result from a loss of sialic acid residues from the mAb carbohydrate epitope. However, binding of HL60var to E- and P-selectin remains sensitive to neuraminidase treatment. Taken together, these data indicate that high levels of surface sLex and/or related epitopes are not essential for interactions with vascular selectins, implying that as yet unidentified sialylated, fucosylated structures serve as physiologically relevant ligands for Eand P-selectin.

DUPLICATE 19 L12 ANSWER 34 OF 43 MEDLINE on STN

ACCESSION NUMBER: DOCUMENT NUMBER:

1999077907 MEDLINE

99077907 PubMed ID: 9858509

TITLE:

Interleukin 12 and interleukin 4 control T cell adhesion to endothelial selectins through opposite effects on alphal,

3-fucosyltransferase VII gene expression.

AUTHOR: CORPORATE SOURCE: Wagers A J; Waters C M; Stoolman L M; Kansas G S

Department of Microbiology-Immunology, Northwestern Medical

School, Chicago, Illinois 60611, USA.

SOURCE:

JOURNAL OF EXPERIMENTAL MEDICINE, (1998 Dec 21) 188 (12)

2225-31.

Journal code: 2985109R. ISSN: 0022-1007.

PUB. COUNTRY:

United States

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT: ENTRY MONTH:

Priority Journals; AIDS

199901

ENTRY DATE:

Entered STN: 19990209

Last Updated on STN: 19990209 Entered Medline: 19990122

ΑВ The alpha1,3-fucosyltransferase, FucT-VII, is crucial for the formation of ligands for all three selectins, and its expression regulates the synthesis of these ligands. Short-term polarized T helper (Th)1, but not Th2 or naive CD4(+) T cells, can home to sites of inflammation, but the molecular basis for this difference has remained unclear. Here we show that naive CD4(+) T cells do not express FucT-VII and fail to bind vascular selectins. We also show that when CD4(+) T cells are activated in the presence of the Th1 polarizing cytokine interleukin (IL)-12, levels of FucT-VII mRNA and binding to E- and P-selectin are significantly augmented. In contrast, activation of CD4(+) T cells in the presence of IL-4, a Th2 polarizing cytokine, inhibited FucT-VII expression and binding to vascular selectins. T cell activation upregulated expression of the Core2 transferase, C2GnT , equivalently regardless of the presence or absence of polarizing cytokines. These data indicate that the selective ability of Th1 cells, as opposed to Th2 cells or naive CD4(+) T cells, to recognize vascular

selectins and home to sites of inflammation is controlled principally by the **expression** of a single gene, FucT-VII.

L12 ANSWER 35 OF 43 MEDLINE on STN DUPLICATE 20

ACCESSION NUMBER: DOCUMENT NUMBER:

1998053883 MEDLINE

98053883 PubMed ID: 9393734

TITLE:

Carcinoma-associated expression of core 2

beta-1,6-N-acetylglucosaminyltransferase gene in human colorectal cancer: role of O-glycans in tumor

progression.

AUTHOR:

Shimodaira K; Nakayama J; Nakamura N; Hasebe O; Katsuyama

T; Fukuda M

CORPORATE SOURCE:

Second Department of Internal Medicine, Shinshu University

School of Medicine, Matsumoto, Japan.

CONTRACT NUMBER:

CA 48737 (NCI)

CA 33000 (NCI)

SOURCE: CANCER RESEARCH, (1997 Dec 1) 57 (23) 5201-6.

Journal code: 2984705R. ISSN: 0008-5472.

PUB. COUNTRY:

United States

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

199803

ENTRY DATE:

Entered STN: 19980312

Last Updated on STN: 19980312 Entered Medline: 19980302

AB Recently, it was demonstrated that an increased level of NeuNAc alpha2-3Gal beta1-4(Fuc alpha1-3)GlcNAc beta-R (sialyl Le(x)) and NeuNAc alpha2-3Gal beta1-3(Fuc alpha1-4)GlcNAc beta-R (sialyl Le(a)) expression on the surface of colorectal cancer cells is positively correlated with progression of the disease. It has not been determined, however, which type of glycans, N- or O-glycans, is more closely associated with progression when cancer cells express those oligosaccharides. To address this problem, we have examined expression of sialyl Le(a) and sialyl Le(x), those oligosaccharides in O-glycans, and core 2 beta-1,6-Nacetylglucosaminyltransferase (C2GnT) transcripts in colorectal cancer specimens from 46 patients and compared those results with clinicopathological variables. C2GnT is a glycosyltransferase that is responsible for the core 2 branch, which is critical for biosynthesis of sialyl Le(a) and sialyl Le(x) in O-glycans. Sialyl Le(a) and sialyl Le(x) were determined by immunohistochemistry, and C2GnT transcripts were detected by reverse transcription-PCR. Sialyl Le(a) or sialyl Le(x) in O-glycans was assessed by combining immunohistochemistry for sialyl Le(a) or sialyl Le(x) with reverse transcription-PCR for C2GnT. Sialyl Le(a), detected on cancer cells in 74% of patients, was well correlated with lymph node metastasis, whereas sialyl Le(a) and sialyl Le(x) in O-glycans, which were specifically detected in cancer tissues of 50 and 61% of patients, respectively, were closely associated with lymphatic and venous invasion. In addition, C2GnT, which was specifically detected in cancer tissues of 63% of patients, was closely correlated with the vessel invasion, as well as depth of tumor invasion. These results strongly suggest that sialyl Le(a) and sialyl Le(x) in O-glycans and C2GnT , expressed in cancer cells, may play important roles in tumor progression through vessel or direct invasion.

L12 ANSWER 36 OF 43 MEDLINE on STN

DUPLICATE 21

ACCESSION NUMBER:

97344242

MEDLINE

DOCUMENT NUMBER:

97344242 PubMed ID: 9224630

TITLE: Expression of stable human O-glycan

core 2 beta-1,6-N-acetylglucosaminyltransferase in Sf9

insect cells.

AUTHOR:

Toki D; Sarkar M; Yip B; Reck F; Joziasse D; Fukuda M;

Schachter H; Brockhausen I

CORPORATE SOURCE: Department of Biochemistry, University of Toronto, Toronto,

Ontario, Canada M5S 1A8.

SOURCE: BIOCHEMICAL JOURNAL, (1997 Jul 1) 325 (Pt 1) 63-9.

Journal code: 2984726R. ISSN: 0264-6021.

PUB. COUNTRY: ENGLAND: United Kingdom

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199708

ENTRY DATE: Entered STN: 19970813

Last Updated on STN: 19970813 Entered Medline: 19970807

AB UDP-GlcNAc:Galbeta1-3GalNAc-R (GlcNAc to GalNAc) beta-1, 6-N-acetylglucosaminyltransferase (C2GnT) catalyses the

formation of O-glycan core 2. Purification and characterization of C2GnT from natural sources has been hampered by the instability of this enzyme. We have been able to prepare a stable partly purified

recombinant human C2GnT by expression

of a truncated form of the enzyme in the baculovirus/Spodoptera frugiperda 9 (Sf9) insect cell system. C2GnT activity was secreted into the Sf9 culture medium (15 pmol/min per microl; approx. 0.2 mg/l) and was stable at 4 degrees C either in solution or after lyophilization. Endoglycosidase H and N-glycanase F treatment of the radiolabelled C2GnT indicated the presence of N-glycans at both potential N-glycosylation sites. The elimination of one or both of the two potential N-glycosylation sites or treatment of the virus-infected insect cells with tunicamycin resulted in loss of enzyme activity due in part to protein degradation.

L12 ANSWER 37 OF 43 MEDLINE on STN DUPLICATE 22

ACCESSION NUMBER:

96216406 MEDLINE

DOCUMENT NUMBER:

96216406 PubMed ID: 8621728

TITLE: Post-translational modifications of recombinant

P-selectin glycoprotein ligand-1 required for binding to P-

and E-selectin.

AUTHOR: Li F; Wilkins P P; Crawley S; Weinstein J; Cummings R D;

McEver R P

CORPORATE SOURCE:

W. K. Warren Medical Research Institute, Department of Medicine, University of Oklahoma Health Sciences Center,

Oklahoma City, 73140, USA.

CONTRACT NUMBER:

P01 HL 54804 (NHLBI)

SOURCE:

JOURNAL OF BIOLOGICAL CHEMISTRY, (1996 Feb 9) 271 (6)

3255-64.

Journal code: 2985121R. ISSN: 0021-9258.

PUB. COUNTRY:

United States

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

199606

ENTRY DATE:

Entered STN: 19960627

Last Updated on STN: 19980206

Entered Medline: 19960619

AB P-selectin glycoprotein ligand-1 (PSGL-1) is a mucin-like ligand for P-and E-selectin on human leukocytes. PSGL-1 requires sialylated, fucosylated O-linked glycans and tyrosine sulfate to bind P-selectin.

Less is known about the determinants that PSGL-1 requires to bind E-selectin. To further define the modifications required for PSGL-1 to bind P- and E-selectin, we transfected Chinese hamster ovary (CHO) cells with cDNAs for PSGL-1 and specific glycosyltransferases. CHO cells synthesize only core 1 O-linked glycans (Galbeta1-3GalNAcalpha1-Se r/Thr); they lack core 2 O-linked glycans (Galbeta1-3(Galbeta1-4GlcNAcbeta1-6)GalNAcalpha1 -Ser/Thr) because they do not express the core 2 beta1 6-N-acetylglucosaminyltransferase (C2GnT). CHO cells also

lack alpha1 3 fucosyltransferase activity. PSGL-1 expressed on transfected CHO cells bound P- and E-selectin only when it was coexpressed with both C2GnT and an alpha1 3 fucosyltransferase (Fuc-TIII, Fuc-TIV, or Fuc-TVII). Chromatography of beta-eliminated O-linked glycans from PSGL-1 co-expressed with C2GnT confirmed synthesis of core 2 structures. Tyrosine residues on PSGL-1 expressed in CHO cells were shown to be sulfated. Phenylalanine replacement of three tyrosines within a consensus sequence for tyrosine sulfation abolished binding to P-selectin but not to E-selectin. These results demonstrate that PSGL-1 requires core 2 O-linked glycans that are sialylated and fucosylated to bind P- and E-selectin. PSGL-1 also requires tyrosine sulfate to bind P-selectin but not E-selectin.

L12 ANSWER 38 OF 43 MEDLINE on STN

DUPLICATE 23

ACCESSION NUMBER:

95173603 MEDLINE

DOCUMENT NUMBER:

PubMed ID: 7869048 95173603

TITLE:

AUTHOR:

Human thymic epithelial cells express

an endogenous lectin, galectin-1, which binds to core 2

O-glycans on thymocytes and T lymphoblastoid cells. Baum L G; Pang M; Perillo N L; Wu T; Delegeane A;

Uittenbogaart C H; Fukuda M; Seilhamer J J

CORPÓRATE SOURCE:

Department of Pathology and Laboratory Medicine, UCLA

School of Medicine 90024.

CONTRACT NUMBER:

AI-07126 (NIAID)

CA-33000 (NCI)

CA-33895 (NCI)

SOURCE:

JOURNAL OF EXPERIMENTAL MEDICINE, (1995 Mar 1) 181 (3)

Journal code: 2985109R. ISSN: 0022-1007.

PUB. COUNTRY:

United States

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

199503

ENTRY DATE:

Entered STN: 19950407

Last Updated on STN: 19950407 Entered Medline: 19950324

AB Thymic epithelial cells play a crucial role in the selection of developing thymocytes. Thymocyte-epithelial cell interactions involve a number of adhesion molecules, including members of the integrin and immunoglobulin superfamilies. We found that human thymic epithelial cells synthesize an endogenous lectin, galectin-1, which binds to oligosaccharide ligands on the surface of thymocytes and T lymphoblastoid cells. Binding of T lymphoblastoid cells to thymic epithelial cells was inhibited by antibody to galectin-1 on the epithelial cells, and by two antibodies, T305 and 2B11, that recognize carbohydrate epitopes on the T cell surface glycoproteins CD43 and CD45, respectively. T lymphoblastoid cells and thymocytes bound recombinant galectin-1, as demonstrated by flow cytometric analysis, and lectin binding was completely inhibited in the presence of lactose. The degree of galectin-1 binding to thymocytes correlated with the maturation stage of the cells, as immature thymocytes bound more galectin-1 than did mature thymocytes. Preferential binding of galectin-1 to immature thymocytes may result from regulated expression of preferred oligosaccharide ligands on those cells, since we found that the epitope recognized by the T305 antibody, the core 2 O-glycan structure on CD43, was expressed on cortical, but not medullary cells. The level of expression of the UDP-GlcNAc: Gal beta 1,3GalNAc-R beta 1, 6GlcNAc transferase (core 2 beta 1, 6 GlcNAc transferase, or C2GnT), which creates the core 2 O-glycan structure, correlated with the glycosylation change between cortical and medullary cells. Expression of mRNA encoding the C2GnT was high in subcapsular and cortical thymocytes and low in

medullary thymocytes, as demonstrated by in situ hybridization. These results suggest that galectin-1 participates in thymocyte-thymic epithelial cell interactions, and that this interaction may be regulated by expression of relevant oligosaccharide ligands on the thymocyte cell surface.

L12 ANSWER 39 OF 43 MEDLINE on STN **DUPLICATE 24**

ACCESSION NUMBER: 96318025 MEDLINE

DOCUMENT NUMBER: 96318025 PubMed ID: 8748164

Isolation and characterization of a pseudogene related to TITLE:

human core 2 beta-1,6-N-acetylglucosaminyl-

transferase.

AUTHOR: Bierhuizen M F; Maemura K; Fukuda M

CORPORATE SOURCE: Glycobiology Program, La Jolla Cancer Research Foundation,

California 92037, USA.

CONTRACT NUMBER: CA33000 (NCI)

CA33895 (NCI)

GLYCOCONJUGATE JOURNAL, (1995 Dec) 12 (6) 857-64. Journal code: 8603310. ISSN: 0282-0080. SOURCE:

PUB. COUNTRY: ENGLAND: United Kingdom

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199611

ENTRY DATE: Entered STN: 19961219

Last Updated on STN: 19961219 Entered Medline: 19961115

In a previous study, we isolated genomic clones encoding core 2 AB beta-1,6-N-acetylglucosaminyltransferase (C2GnT) and blood group IGnT and proposed that these two genes were produced from a common ancestral gene by duplication, diversion and intron insertion. In the present study, we have isolated a pseudogene which is highly related to the gene of C2GnT. The sequence analysis of this pseudogene indicated that the pseudogene was produced by duplication of a common precursor gene for C2GnT. These results taken together strongly suggest that the ancestral gene was first duplicated and one of the duplicated genes directly evolved into the IGnT gene. The other duplicated gene was further duplicated to produce the C2GnT gene and the pseudogene.

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on STN **DUPLICATE 25**

ACCESSION NUMBER: 95225818 EMBASE

DOCUMENT NUMBER: 1995225818

TITLE: Genomic organization of core 2 and I branching

.beta.-1,6-N- acetylglucosaminyltransferases. Implication

for evolution of the .beta.-1,6-N-

acetylglucosaminyltransferase gene family.

AUTHOR: Bierhuizen M.F.A.; Maemura K.; Kudo S.; Fukuda M.

CORPORATE SOURCE: La Jolla Cancer Research Foundation, 10901 North Torrey

Pines Road, San Diego, CA 92037, United States

SOURCE: Glycobiology, (1995) 5/4 (417-425).

ISSN: 0959-6658 CODEN: GLYCE3

COUNTRY: United Kingdom DOCUMENT TYPE: Journal; Article

FILE SEGMENT: Clinical Biochemistry 029

LANGUAGE: English SUMMARY LANGUAGE: English

Two human .beta.-1,6-N-acetylglucosaminyltransferases forming the core 2 O- glycan branch, C2GnT and the I antigen IGnT, are homologous to each other in three regions of the catalytic domain (A, B, C) and their genes reside at the same locus, chromosome 9, band q21 (Bierhuizen, M.F.A., Mattei, M.-G. and Fukuda, M., Genis Dev., 7, 468-478, 1993). In order to investigate how these two enzymes are related at the

genomic level, and how this gene family evolved, we have elucidated their genomic structures. It was found that C2GnT is coded by two exons, of which the second exon encodes the whole translation product. In contrast, the complete coding sequence for IGnT is divided over three exons. Importantly, the highly homologous region B is encoded entirely by exon 2 in the C2GnT gene, while the same region is split between exons 1 and 2 in the IGnT gene. The other highly homologous regions, A and C, are also encoded by exon 2 in the C2GnT gene, while they are encoded by exon 1 and exon 3, respectively, in the IGnT gene. These results strongly suggest that the common ancestral gene was first duplicated and then each duplicated gene evolved into the C2GnT or IGnT gene by intron insertion and divergence following the duplication. The sequences upstream from the transcription initiation sites of the C2GnT and IGnT genes have promoter activity and contain TATA-like sequences. In addition, the promoter sequence of the C2GnT gene contains potential binding sites for a variety of transcription factors, including NF-IL6, GATA-3 and TCF-1, which are specifically active in T lymphocytes and during inflammation. The results are consistent with the fact that C2GnT is highly expressed in activated T lymphocytes and myeloid cells.

L12 ANSWER 41 OF 43 BIOTECHDS COPYRIGHT 2004 THOMSON DERWENT/ISI on STN

ACCESSION NUMBER: 1994-07116 BIOTECHDS

TITLE: New beta-1,6-N-acetylglucosaminyltransferase and acceptor;

human recombinant enzyme production by vector plasmid pcDNAI-C2GnT expression

in CHO-Py-leu cell culture

PATENT ASSIGNEE: La-Jolla-Cancer-Res.Found.

PATENT INFO: EP 590747 6 Apr 1994

APPLICATION INFO: EP 1993-250268 29 Sep 1993 PRIORITY INFO: US 1992-955041 1 Oct 1992

DOCUMENT TYPE: Patent LANGUAGE: English

OTHER SOURCE: WPI: 1994-111195 [14]

AB **Human recombinant** purified beta-1,6-N-

acetylglucosaminyltransferase (I) (EC-2.4.1.150) or a (I) fragment is claimed. (I) has activity of UDP-GlcNAc:Gal-beta-1-3GalNAc (GlcNAc to GalNAc) beta-1-6 N-acetylglucosaminyltransferase. (I) preferably has mol.wt. about 50,000. Also claimed are: (1) nucleic acid (II) encoding (I) or a (I) fragment (DNA sequence disclosed); (2) a vector containing (II), especially plasmid pcDNAI-C2GnT; (3) a host cell containing the vector; (4) a purified human protein or a protein fragment that is an acceptor molecule (III) which is acted upon by (I) having activity which forms core 2 oligosaccharide structures in O-glycans; (5) nucleic acid encoding (III); (6) a vector containing the nucleic acid of (5), especially vector plasmid pcDSR-alpha-leu; (7) a host cell containing the vector of (6); (8) a method for obtaining from a cell line recombinant (I) or recombinant (III) involving transfecting a cell line (preferably CHO-Py-leu) with a vector and screening the gene bank for (I) or (III) expression; and (9) a method for isolating (I) from a host cell. (I) is useful as a tumor marker and (III) can be used for antibody production. (34pp)

L12 ANSWER 42 OF 43 MEDLINE on STN DUPLICATE 27

ACCESSION NUMBER: 941408

94140881 MEDLINE

94140881 PubMed ID: 8308016

TITLE:

AUTHOR:

Expression of a differentiation antigen and

poly-N-acetyllactosaminyl O-glycans directed by a

cloned core 2 beta-1,6-N-

acetylglucosaminyltransferase. Bierhuizen M F; Maemura K; Fukuda M

CORPORATE SOURCE:

DOCUMENT NUMBER:

Glycobiology Program, La Jolla Cancer Research Foundation,

California 92037.

CONTRACT NUMBER:

· CA33000 (NCI)

CA33895 (NCI)

SOURCE: JOURNAL OF BIOLOGICAL CHEMISTRY, (1994 Feb 11) 269 (6)

4473-9.

Journal code: 2985121R. ISSN: 0021-9258.

PUB. COUNTRY:

United States

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

199403

ENTRY DATE:

Entered STN: 19940330

Last Updated on STN: 19940330 Entered Medline: 19940317

AΒ Chinese hamster ovary (CHO) cells do not contain detectable amounts of core 2 beta-1,6-N-acetylglucosaminyltransferase, C2GnT, and thus lack various modifications in their branched O-linked oligosaccharides. In the present study, the O-linked oligosaccharides and the occurrence of a differentiation antigen were analyzed in CHO cells stably transfected with cDNA encoding human leukosialin alone (CHO-leu) or with cDNAs encoding both leukosialin and C2GnT (CHO-leu.C2GnT). The analysis of O-glycans, released from [3H]glucosamine-labeled cells, revealed that CHO-leu cells synthesize O-glycans with a Gal beta 1-->3GalNAc backbone, whereas CHO-leu.C2GnT cells synthesize in addition O-glycans with a Gal beta 1-->3 (Gal beta 1-->4GlcNAc beta 1-->6) GalNAc backbone. Moreover, CHO-leu.C2GnT cells express poly-N-acetyllactosaminyl extensions from the GlcNAc beta 1-->6 branch in O-glycans, while CHO-leu cells express no detectable amount of poly-N-acetyllactosaminyl O-glycans. It was also demonstrated that leukosialin in CHO-leu. C2GnT cells is recognized by the T305 monoclonal antibody, while the same antibody did not react at all with CHO-leu cells. In addition, the transient expression cloning scheme using the T305 monoclonal antibody as a selectin marker and COS-1 cells, which endogenously express C2GnT as recipient cells, resulted in the isolation of cDNA encoding leukosialin. These results indicate that C2GnT determines the expression of poly-Nacetyllactosamines in O-glycans and together with leukosialin, an onco-differentiation antigen recognized by the T305 antibody.

L12 ANSWER 43 OF 43 SCISEARCH COPYRIGHT 2004 THOMSON ISI on STN

ACCESSION NUMBER:

95:129212 SCISEARCH

THE GENUINE ARTICLE: OF341

TITLE:

DIFFERENTIAL REGULATION OF CD43 GLYCOFORMS ON CD4(+) AND

CD8(+) T-LYMPHOCYTES IN GRAFT-VERSUS-HOST DISEASE

AUTHOR: ELLIES L G (Reprint); JONES A T; WILLIAMS M J; ZILTENER H

CORPORATE SOURCE:

UNIV BRITISH COLUMBIA, BIOMED RES CTR, 2222 HLTH SCI MALL,

VANCOUVER, BC V6T 1Z3, CANADA (Reprint); UNIV BRITISH COLUMBIA, DEPT PATHOL & LAB MED, VANCOUVER, BC V6T 1Z3,

CANADA

COUNTRY OF AUTHOR:

CANADA

SOURCE:

GLYCOBIOLOGY, (DEC 1994) Vol. 4, No. 6, pp. 885-893.

ISSN: 0959-6658. Article; Journal

DOCUMENT TYPE: FILE SEGMENT:

LIFE

LANGUAGE:

ENGLISH

REFERENCE COUNT: 53

ABSTRACT IS AVAILABLE IN THE ALL AND IALL FORMATS

AB Two distinct T-cell glycoforms of CD43 result from differential glycosylation of a single gene product in vivo. The 115 kDa glycoform tarries mainly tetrasaccharides and is a pan T-cell marker, whereas the 130 kDa glycoform carries mainly hexasaccharides and is associated with T-cell activation, CD43 has been shown to play a role both in enhancing and inhibiting cell adhesion; however, the function of the individual glycoforms is unknown. We have examined the distribution and regulation of

the CD43 glycoforms in a murine model of acute graft-versus-host disease (GVHD) using monoclonal antibodies (mAbs) S7 and 1B11 specific for the 115 and 130 kDa CD43 glycoforms, respectively, An increase in T-lymphocyte CD43 130 kDa expression occurred during GVHD from day 4 onwards and coincided with splenomegaly and upregulation of the beta 1-6GlcNAc transferase (C2GnT), the key enzyme responsible for the addition of complex O-glycan branching to CD43. When T-lymphocyte subsets were examined for CD43 expression, we found that in GVHD, both CD43 glycoforms were upregulated on CD4(+) T cells. However, in CD8(+) T cells, CD43 115 kDa was downregulated while CD43 130 kDa was dramatically upregulated, such that two distinct CD8(+)1B11(+) T-cell subsets were observed. These data demonstrate differential expression of the CD43 glycoforms in both resting and activated CD4(+) and CD8(+)T cells, and suggest that glycosylation differences between the CD43 glycoforms may reflect participation in the different functions of these T-cell subsets in immune disorders in vivo.

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L1
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L2
               0 S ACETYLKGLYCOSAMINE
1.3
          36991 S ACETYLGLUCOSAMINE
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L4
L5
              4 DUP REM L1 (6 DUPLICATES REMOVED)
L6
            257 S "C2GNT"
           5011 S "N-ACETYLGLUCOSAMINYLTRANSFERASE?"
L7
           5065 S L6 OR L7
L8
        6346882 S CLON? OR EXPRESS? OR RECOMBINANT
L9
            233 S L6 AND L9
L10
L11
            146 S HUMAN AND L10
            43 DUP REM L11 (103 DUPLICATES REMOVED)
L12
                E SCHWIENTEK T/AU
            116 S E3-E4
L13
                E CLAUSEN H/AU
            804 S E3
L14
            868 S L13 OR L14
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PROCESSING COMPLETED FOR L16
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    ANSWER 1 OF 1
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L17
AN
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     99143102
                PubMed ID: 9988682
TI
     Control of O-glycan branch formation. Molecular cloning of human
     cDNA encoding a novel beta1,6-N-acetylglucosaminyltransferase forming core
AU
     Schwientek T; Nomoto M; Levery S B; Merkx G; van Kessel A G;
     Bennett E P; Hollingsworth M A; Clausen H
CS
     School of Dentistry, University of Copenhagen, Norre Alle 20, 2200
     Copenhagen N, Denmark.
     1 RO1 CA66234 (NCI)
NC
     1RO1 CA66234 (NCI)
     5 P41 RR05351 (NCRR)
     JOURNAL OF BIOLOGICAL CHEMISTRY, (1999 Feb 19) 274 (8) 4504-12.
     Journal code: 2985121R. ISSN: 0021-9258.
CY
     United States
DT
     Journal; Article; (JOURNAL ARTICLE)
LA
     English
FS
     Priority Journals
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     GENBANK-AF038650
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     199903
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     Entered STN: 19990326
    Last Updated on STN: 20000303
    Entered Medline: 19990318
    A novel human UDP-GlcNAc:Gal/GlcNAcbeta1-3GalNAcalpha beta1,
AB
     6GlcNAc-transferase, designated C2/4GnT, was identified by BLAST analysis
     of expressed sequence tags. The sequence of C2/4GnT encoded a
    putative type II transmembrane protein with significant sequence
     similarity to human C2GnT and IGnT. Expression of the
     secreted form of C2/4GnT in insect cells showed that the gene product had
    UDP-N-acetyl-alpha-D-glucosamine:acceptor beta1, 6-N-
     acetylglucosaminyltransferase (beta1,6GlcNAc-transferase) activity.
    Analysis of substrate specificity revealed that the enzyme catalyzed
     O-glycan branch formation of the core 2 and core 4 type. NMR analyses of
     the product formed with core 3-para-nitrophenyl confirmed the product core
     4-para-nitrophenyl. The coding region of C2/4GnT was contained in a
     single exon and located to chromosome 15q21.3. Northern analysis revealed
     a restricted expression pattern of C2/4GnT mainly in colon,
    kidney, pancreas, and small intestine. No expression of C2/4GnT
    was detected in brain, heart, liver, ovary, placenta, spleen, thymus, and
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peripheral blood leukocytes. The **expression** of core 2 O-glycans has been correlated with cell differentiation processes and cancer. The results confirm the predicted existence of a beta1,6GlcNAc-transferase that functions in both core 2 and core 4 O-glycan branch formation. The redundancy in beta1,6GlcNAc-transferases capable of forming core 2 O-glycans is important for understanding the mechanisms leading to specific changes in core 2 branching during cell development and malignant transformation.

Check Tags: Human; Male; Support, Non-U.S. Gov't; Support, U.S. Gov't, P.H.S.

Amino Acid Sequence

Base Sequence

Chromosomes, Human, Pair 15

Cloning, Molecular

DNA, Complementary

Magnetic Resonance Spectroscopy

Molecular Sequence Data

N-Acetylglucosaminyltransferases: CH, chemistry

*N-Acetylglucosaminyltransferases: GE, genetics

*Polysaccharides: CH, chemistry

Protein Conformation

Sequence Homology, Amino Acid

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 L_5

L6

L8

L10

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(FILE 'HOME' ENTERED AT 14:09:33 ON 30 JAN 2004)

FILE 'MEDLINE, EMBASE, BIOSIS, BIOTECHDS, SCISEARCH, HCAPLUS, NTIS, LIFESCI' ENTERED AT 14:11:05 ON 30 JAN 2004

L1 10 S "C2GNT3"

L2 0 S ACETYLKGLYCOSAMINE

L3 36991 S ACETYLGLUCOSAMINE

L4 0 S "N-ACETYLGLUCVOSAM

0 S "N-ACETYLGLUCVOSAMINETRANSFERASE?"

4 DUP REM L1 (6 DUPLICATES REMOVED)

257 S "C2GNT"

L7 5011 S "N-ACETYLGLUCOSAMINYLTRANSFERASE?"

5065 S L6 OR L7

L9 6346882 S CLON? OR EXPRESS? OR RECOMBINANT

233 S L6 AND L9

L11 146 S HUMAN AND L10

L12 43 DUP REM L11 (103 DUPLICATES REMOVED)

E SCHWIENTEK T/AU

L13 116 S E3-E4

E CLAUSEN H/AU

L14 804 S E3

L15 868 S L13 OR L14

L16 6 S L10 AND L15

L17 1 DUP REM L16 (5 DUPLICATES REMOVED)

	Issue Date	Pages	Document ID	Title
1	20040129	97	US 20040018590 A1	Combinatorial DNA library for producing modified N-glycans in lower eukaryotes
2	20040115	60	US 20040009477 A1	Methods for producing libraries of expressible gene sequences
3	20040108	194	US 20040005633 A1	Methods and apparatuses for gel-free qualitative and quantitative proteome analysis, and uses therefore
4	20031211	119	US 20030228664 A1	Secreted and transmembrane polypeptides and nucleic acids encoding the same
5	20031204	106	US 20030224411 A1	Genes that are up- or down-regulated during differentiation of human embryonic stem cells
6	20031127	22	US 20030219830 A1	Methods of evaluating glycomolecules for enhanced activities
7	20031002	125	US 20030186364 A1	Staphylococcus aureus genes and polypeptides
8	20030925	35	US 20030180778 Al	UDP-N-acetylglucosamine: galactose-beta1,3-N-acetylgal actosamine-alpha-R/ (GlcNAc to GalNAc) beta1,6-N-acetylglucosaminylt ransferase, C2GnT3
9	20030918	142	US 20030175902 A1	Methods for producing hyaluronan in a recombinant host cell
10	20030911	121	US 20030171275 A1	Transporters and ion channels
11	20030911	125	US 20030170864 A1	Secreted and transmembrane polypeptides and nucleic acids encoding the same

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	Issue Date	Pages	Document ID	Title
12	20030807	64	US 20030148460 A1	Phosphodiester alpha-GlcNAcase of the lysosomal targeting pathway
13	20030717	102	US 20030134302 A1	Libraries of expressible gene sequences
14	20030612	81	US 20030108872 A1	Genomics-assisted rapid identification of targets
15	20030529	53	US 20030099967 A1	Heparin/heparosan synthase from P. multocida and methods of making and using same
16	20030522	22	US 20030096281 A1	Methods of making glycomolecules with enhanced activities and uses thereof
17	20030508	87	US 20030087818 A1	Compositions and methods for the therapy and diagnosis of colon cancer
18	20030424	125	US 20030077657 Al	Secreted and transmembrane polypeptides and nucleic acids encoding the same
19	20030424	166	US 20030077568 A1	Methods of diagnosis of colorectal cancer, compositions and methods of screening for colorectal cancer modulators
20	20030417	102	US 20030073163 A1	Libraries of expressible gene sequences
21	20030320	: < 5	US 20030054525 A1	UDP-N-acetylglucosamine: galactose-beta1,3-N-acetylgal actosamine-alpha-R / (GlcNAc to GalNAc) beta1,6-N-acetylglucosaminylt ransferase, C2GnT3

	Issue Date	Pages	Document ID	Title
22	20030320	118	US 20030054447 A1	Secreted and transmembrane polypeptides and nucleic acids encoding the same
23	20030213	124	US 20030032060 A1	Secreted and transmembrane polypeptides and nucleic acids encoding the same
24	20030206		US 20030027256 A1	Secreted and transmembrane polypeptides and nucleic acids encoding the same
25	20021219		US 20020192678 A1	Genes expressed in senescence
26	20021219		US 20020192668	Secreted and transmembrane polypeptides and nucleic acids encoding the same
27	20021205		US 20020182618 A1	Secreted and transmembrane polypeptides and nucleic acids encoding the same
28	20021128		US 20020177182 A1	Methods for the identification of antimicrobial compounds
29	20021107	4h :	US 20020164748 A1	Glycosyl sulfotransferase-3

	Issue Date	Pages	Document ID	Title
30	20021107		US 20020164646 A1	Secreted and transmembrane polypeptides and nucleic acids encoding the same
31	20021031		US 20020160392 A1	Secreted and transmembrane polypeptides and nucleic acids encoding the same
32	20021017	298	US 20020151681 A1	Nucleic acids, proteins and antibodies
33	20021017		US 20020150981 A1	Methods for producing highly phosphorylated lysosomal hydrolases
34	20021003	88	US 20020142386 A1	Engineering intracellular sialylation pathways
35	20020912		US 20020128224 A1	Compositions and methods for the treatment of glaucoma or ocular hypertension
36	20020725		US 20020098507 A1	Secreted and transmembrane polypeptides and nucleic acids encoding the same
37	20020725		US 20020098506 A1	Secreted and transmembrane polypeptides and nucleic acids encoding the same
38	20020725		US 20020098505 A1	Secreted and transmembrane polypeptides and nucleic acids encoding the same

	Issue Date	Pages	Document ID	Title
39	20020627		US 20020081656 A1	UDP-N-acetylglucosamine: Galactose-beta1,3-N-acetylgal actosamine-alpha-R / N-acetylglucosamine-beta1,3,- N-acetylgalactosamine-alpha-R (GlcNAc to GalNAc) beta1,6-N-acetylglucosaminylt ransferase, C2/4GnT
40	20020530		US 20020064816 A1	Moss genes from physcomitrella patens encoding proteins involved in the synthesis of carbohydrates
41	20020516		US 20020059659 A1	DNA shuffling to produce herbicide selective crops
42	20020516		US 20020058249 A1	DNA SHUFFLING TO PRODUCE HERBICIDE SELECTIVE CROPS
43	20020228		US 20020025550 A1	Methods for producing highly phosphorylated lysosomal hydrolases
44	20011213	27	US 20010051370 A1	Glycosyl sulfotransferase-3
45	20031230		US 6670165 B2	Methods for producing highly phosphorylated lysosomal hydrolases
46	20031104		US 6642038 B1	GlcNAc phosphotransferase of the lysosomal targeting pathway
47	20031021		US 6635461 B1	UDP-N-acetylglucosamine: galactosebeta.1, 3-N-acetylgalactosaminealph aR/(GlcNAc to GalNAc) .beta.1,6-N-acetylglucosaminy ltransferase, C2GnT3
48	20030805		US 6602693 B1	Gene encoding hyaluronan synthase
49	20030722		US 6596523 B1	.alpha.,2,8-sialyltransferase
50	20030513		US 6562958 B1	Nucleic acid and amino acid sequences relating to Acinetobacter baumannii for diagnostics and therapeutics

	Issue Date	Pages	Document	ID	Title
51	20030325		US 6537785	B1	Methods of treating lysosomal storage diseases
52	20030304		US 6528289	B1	Nucleotide sequence of the Haemophilus influenzae Rd genome, fragments thereof, and uses thereof
53	20030114		US 6506581	B1	Nucleotide sequence of the Haemophilus influenzae Rd genome, fragments thereof, and uses thereof
54	20021210		US 6492150	B1	Gene encoding hyaluronan synthase
55	20020611	123	US 6403337	B1	Staphylococcus aureus genes and polypeptides
56	20020521		US 6391614	B1	Auxiliary gene and protein of methicillin resistant bacteria and antagonists thereof
57	20020402	38	US 6365365	B1	Method of determining whether an agent modulates glycosyl sulfotransferase-3
58	20020326		US 6361995	B1	Protection of pancreatic .betacells during islet isolation and assessment of islet viability and candidate diabetes drugs after islet isolation
59	20020312		US 6356845	B1	Crystallization and structure determination of Staphylococcus aureus UDP'N-acetylenolpyruvylglucos amine reductase (S. aureus MurB)
60	20020101	227 [.]	US 6335170	B1	Gene expression in bladder tumors

	Issue Date	Pages	Document	ID	Title
61	20010724	27	US 6265192	B1	Glycosly sulfortransferase-3
62	20010626		US 6251647	B1	Auxiliary genes and proteins of methicillin resistant bacteria and antagonists thereof
63	20010508		US 6228627	В1	Mur A-1 from Streptococcus pneumoniae
64	20010508		US 6228612	B1	Compounds
65	20010320		US 6204042	B1	GlmU
66	20001024	*	US 6136580	· A	.beta1-6-N-acetylglucosamin yltransferase that forms core 2, core 4 and I branches
67	20000801		US 6096512	A	Cloned DNA encoding a UDP-GalNAc: Polypeptide, N-acetylgalactosaminyltransfe rase
68	20000328		US 6043071	A	GlmU

	Issue Date	Pages	Document	ID	Title
69	20000118	64	US 6015701	Α	N-acetylglucosaminyltransfera se V proteins and coding sequences
70	19991116		US 5985643	A	Auxiliary gene and protein of methicillin resistant bacteria and antagonists thereof
71	19990608		US 5910570	Α	Cloned DNA encoding a UDP-GalNAc: polypeptide N-acetylgalactosaminy-ltransf erase
72	19981110		US 5834284	A	N-acetylglucosaminyl transferase gene coding therefor and process for production thereof
73	19980616		US 5766910	A	Expression of the developmental I antigen by a cloned human cDNA encoding a member of a beta-1, 6-N-acetylglucosaminyltransfr ase gene family
74	19980324		US 5731420	A	Antibodies to human I-branching beta-1,6-N-acetylglucosaminyl transferase
75	19971104		US 5684134	A	Antibody specific for beta.1.fwdarw.6 N-acetylglucosamininyltransfe rase
76	19970819		US 5658778	A	.beta.1-6 N-acetylglucosaminyl, transferase, its acceptor molecule, leukosialin, and a method for cloning proteins having enzymatic activity
77 .	19970429	·	US 5624832	Α	.beta.1 6 N-acetylglucosaminyltransfera se, its acceptor molecule, leukosialin, and a method for cloning proteins having enzymatic activity
78	19960116		US 5484590	Α	Expression of the developmental I antigen by a cloned human cDNA encoding a member of a .beta1,6-N-acetylglucosamin yltransferase gene family

	Issue Date	Pages	Document II	D	Title
79	19941101		US 5360733 A		Human .beta.1-6 n-acetylglucosaminyl transferase

	Issue Date	Pages	Document ID	Title
1	20030925	35	US 20030180778 A1	to GalNAc) beta1,6-N-acetylglucosaminylt ransferase, C2GnT3
2	20030501	78	US 20030082511 A1	Identification of modulatory molecules using inducible promoters
3	20030320	35	US 20030054525 A1	UDP-N-acetylglucosamine: galactose-beta1,3-N-acetylgal actosamine-alpha-R / (GlcNAc to GalNAc) beta1,6-N-acetylglucosaminylt ransferase, C2GnT3
4	20021107	36	US 20020164748 A1	Glycosyl sulfotransferase-3
5	20020627	25	US 20020081656 A1	UDP-N-acetylglucosamine: Galactose-beta1,3-N-acetylgal actosamine-alpha-R / N-acetylglucosamine-beta1,3,- N-acetylgalactosamine-alpha-R (GlcNAc to GalNAc) beta1,6-N-acetylglucosaminylt ransferase, C2/4GnT
5	20011213	27	US 20010051370 A1	Glycosyl sulfotransferase-3
7	20031021	34	US 6635461 B1	UDP-N-acetylglucosamine: galactosebeta.1, 3-N-acetylgalactosaminealph aR/(GlcNAc to GalNAc) .beta.1,6-N-acetylglucosaminy ltransferase, C2GnT3
8	20020402	38	US 6365365 B1	Method of determining whether an agent modulates glycosyl sulfotransferase-3
)	20010724	27	US 6265192 B1	Glycosly sulfortransferase-3

	Issue Date	Pages	Document	ID	Title
10	20001024	30	US 6136580	Α	.beta1-6-N-acetylglucosamin yltransferase that forms core 2, core 4 and I branches
11	20000926	37	US 6124267	A	O-glycan inhibitors of selectin mediated inflammation derived from PSGL-1
12	19980616	31	US 5766910	Α	Expression of the developmental I antigen by a cloned human cDNA encoding a member of a beta-1, 6-N-acetylglucosaminyltransfrase gene family
13	19980324	30	US 5731420	Α	Antibodies to human I-branching beta-1,6-N-acetylglucosaminyl transferase
14	19971104	26	US 5684134	A	Antibody specific for .beta.1.fwdarw.6 N-acetylglucosamininyltransfe rase
15	19970819	24	US 5658778	A	.beta.1-6 N-acetylglucosaminyl, transferase, its acceptor molecule, leukosialin, and a method for cloning proteins having enzymatic activity
16	19970429	26	US 5624832	Α	.beta.1 6 N-acetylglucosaminyltransfera se, its acceptor molecule, leukosialin, and a method for cloning proteins having enzymatic activity
17	19970408	23	US 5619726	A	Apparatus and method for performing arbitration and data transfer over multiple buses
18	19960116	31	US 5484590	A	Expression of the developmental I antigen by a cloned human cDNA encoding a member of a .beta1,6-N-acetylglucosamin yltransferase gene family

	Issue Date	Pages	Document ID	Title
19				Human .beta.1-6 n-acetylglucosaminyl transferase

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	L #	Hits	Search Text
1	L2	0	"glucosaminetransferase\$2"
2	L3	0	glucosaminyltransferase\$2"
3	L4	215	"UDP-N-acetylglucosamine"
4	L5	230	l1 or l4
5	L6	595126	clon\$3 or express\$3 or recombinant
6	L7	79	15 same 16
7	L1	19	"c2GnT"
8	L8	6	schwientek.in.
9	L9	672	clausen.in.
10	L10	674	18 or 19
11	L11	4	110 and 11